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(54) Title: CHLAMYDIA TRACHOMATIS GENOMIC SEQUENCE AND POLYPEPTIDES, FRAGMENTS THEREOF AND USES THEREOF, IN PARTICULAR FOR THE DIAGNOSIS, PREVENTION AND TREATMENT OF INFECTION

(57) Abstract

The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of Chlamydia trachomatis, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences, as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the Chlamydia trachomatis genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing Chlamydia trachomatis infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular Chlamydia trachomatis, infections.

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CHLAMYDIA TRACHOMATIS GENOMIC SEQUENCE AND POLYPEPTIDES, FRAGMENTS THEREOF AND USES THEREOF, IN PARTICULAR FOR THE DIAGNOSIS, PREVENTION AND TREATMENT OF INFECTION

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The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of *Chlamydia trachomatis*, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the *Chlamydia trachomatis* genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing *Chlamydia trachomatis* infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular *Chlamydia trachomatis*, infections.

The genus Chlamydia is composed of four species: Chlamydia psittaci, Chlamydia 20 pecorum, Chlamydia pneumoniae and Chlamydia trachomatis.

Chlamydia psittaci comprises numerous species, whose hosts are terrestrial vertebrate animals as well as birds and occasionally humans;

Chlamydia pecorum is a pathogen of ruminants;

Chlamydia pneumoniae is responsible for pneumopathies, for sinusitis and for arterial impairments in humans;

Chlamydia trachomatis (Ct) is responsible for a large number of human diseases:

- eye diseases: conventional trachoma, nonendemic trachoma, paratrachoma, inclusion conjunctivitis in neonates and in adults;
- genital diseases: nongonococcal uretritis, epididymitis, cervicitis, salpingitis, perihepatitis and bartholinitis as well as pneumopathy in breast-feeding infants;
 - systemic diseases: venereal lymphogranulomatosis (LGV).

These diseases affect a very large number of women and men [more than 600 million individuals are trachoma carriers and there are more than 90 million cases of genital *Chlamydia* infections] worldwide. Accordingly, basic and applied research which makes it possible to understand the physiopathology linked to this bacterium is very important for public health. (Raulston JE., 1995; Hackstadt T. et al., 1996).

Eye impairments due to Chlamydia trachomatis cause trachoma and inclusion

conjunctivitis. Trachoma is a chronic conjunctivitis. It is the major cause of curable eye diseases leading to blindness. It is estimated that 20 million cases of loss of sight are due to it worldwide. Moreover, inclusion conjunctivitis is an eye inflammation which is caused by *Chlamydia trachomatis* and is transmitted by the venereal route. Inclusion conjunctivitis affects adults and neonates exposed to genital secretions.

Two types of eye disease caused by agents of the species Chlamydia trachomatis can be distinguished. The conventional trachomatous disease is found in endemic regions; transmission occurs from eye to eye and through the hands, or it can be passed on by flies. In nonendemic regions, transmission occurs through the genital apparatus; it usually only causes conjunctivitis, most often 10 without associated keratitis; it is rare for a pannus or for scars similar to those in trachoma to develop. This conjunctival impairment is called paratrachoma to differentiate it from the conventional endemic trachoma which is transmitted by the ocular route. The seriousness and the number of cases of trachoma have decreased over the last forty years. This is related to the improvement in hygiene and living conditions. However, trachoma remains the principal cause of avoidable blindness in Africa, in 15 the Middle East and in some regions of Asia. The transmission of the endemic disease occurs in particular through close personal contact, in regions where a secondary exposure exists in a repeated form. Often, the infection is also latent. In some industrialized countries, such as the United States, a mild form of trachoma still exists in some ethnic groups. Sometimes, a tardive trachoma may be found following an immunosuppressive treatment. The eye impairments caused by Chlamydia trachomatis, such as inclusion conjunctivitis and paratrachoma, are also a complication due to a common venereal infection. These infections are not very frequent; they occur most often in young adults. The eye impairments in neonates are produced during the passage through the maternal genital routes during childbirth. Theoretically, endemic trachoma and inclusion conjunctivitis in adults appear in the form of conjunctivitis, the latter being characterized by the presence of lymphoid follicles. In regions where 25 the endemic disease is serious, the disease often starts before the age of 2 years and reinfection is frequent. Superficial neovascularization is added, in this case, to leukocytic infiltration. The conjunctival scars will then cause trichiasis and entropion. The eroded comea will become a carrier of a corneal ulcer of bacterial origin. The scar on the cornea causes blindness. Impairment of the lachrymal glands gives a picture of dryness of the comea. Xerosis becomes complicated with 30 secondary bacterial ulcer. In regions where trachoma is endemic, the infectious process disappears towards the age of fifteen. The scars then progress to blindness, which affects almost exclusively adults. In regions where exposure is lower, the infectious process is, in this case, less rapid and adults are carriers of a chronic disease.

Positive diagnosis of trachoma can be most often established by clinical observation:

35 lymphoid follicles are visible on the upper tarsal conjuctiva; conjunctival scar is typical. Vascular

pannus exists. In endemic regions, clinical diagnosis is often sufficient. However, isolated cases of inclusion conjunctivitis must be the subject of a differential diagnosis, in particular to distinguish viral conjunctivitis.

Public health measures against the endemic form of the disease provide for mass treatments with tetracycline or erythromycin collyria of all children. The treatment may also provide for surgical correction of the lesions. The other conjunctival impairments respond well to general treatments with tetracyclines or erythromycin. The prevention of trachomatous disease by health measures and by improving living standards is sufficient. Furthermore, to avoid the spread of trachoma, antibiotic collyria may be used.

The role of *Chlamydia trachomatis* in a number of genital impairments has been demonstrated over the last three decades. *Chlamydia trachomatis* is responsible in this case for a pathology which may be superposed on the impairments observed with *Neisseria gonorrhoeae*. The pathologies for which *Chlamydia trachomatis* may be responsible at the genital level are acquired by the venereal route and are a major source of sexually transmitted diseases.

The epidemiology of *Chlamydia trachomatis* genital infections shows each year more than 4 million new cases in the United States, and more than 3 million new cases in Europe. Like the other venereal infections, *Chlamydia trachomatis* affects young subjects. There is a direct relationship between the number of sexual partners and the frequency of the disease. For example, the frequency of *Chlamydia trachomatis* appears to be five to ten times higher than that of *Neisseria gonorrhoeae* in pregnant women. The *Chlamydia trachomatis* infection is probably more discreet than its *Neisseria gonorrhoeae* homologue. This relative clinical silence, estimated in women at 50% or even 70% of infections, explains why the total morbidity of *Chlamydia trachomatis* conditions is high. Diagnosis must therefore be requested in patients who are sometimes asymptomatic carriers of infection.

Chlamydia trachomatis is responsible for nearly 30% of nongonococcal urethritis, or NGU. Chlamydia trachomatis urethritis may be discreet, the disease then progresses to a certain form of chronicity. The diagnosis will, like for the other clinical forms of the disease, be called into play later.

Chlamydia trachomatis is a cause of epididymitis in humans during a period of sexual activity. The bacterium may be found in the urethra, urine, sperm or even a sample collected by aspiration from the epididymis. It is in particular found in humans under 35 years of age. A discharge from the urethra which is associated with the disease suggests the diagnosis of a Chlamydia condition or sometimes a gonococcal condition.

Untreated Reiter's syndrome, if accompanied by urethritis, evokes a Chlamydia trachomatis condition.

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gonorrhoea (or have had contact), 10% to 20% of women having a venereal origin, 5% of women consulting having no particular origin.

The cervix is often normal during a Chlamydia trachomatis infection. However, a hypertrophic cervical erythema will cause such an infection to be suspected. Chlamydia trachomatis is responsible for an endocervicitis whereas viral impairments result in exocervicitis. A nongonococcal endocervicitis requires treating the patient and partners with tetracyclines.

Chlamydia trachomatis is responsible for a large number of acute salpingites. The picture is often complicated by an acute peritonitis or even a perihepatitis.

In case of pregnancy, the risk is first that of infection of the neonate at birth. However, the risk of postpartum complications exists (endometritis or salpingitis).

The reference method for the diagnosis of *Chlamydia trachomatis* is the isolation of the bacterium on cell culture. For all infections, the sample collection should make it possible to obtain a suitable sample with the aid of a swab. This sample should be transported to a laboratory under excellent conditions; in particular, the cold chain must absolutely be maintained. The placing in cell culture on mouse fibroblasts will be carried out by people having specific skills. The distinction of *Chlamydia trachomatis* with labelled antibodies and the observation of cell cultures under a microscope will take place two days after placing in culture. Provided these imperatives are observed, cell culture is a reliable technique. However, the constraints linked to this technique are many: not only must the laboratory be equipped for the cell culture, but, furthermore, highly competent staff must take care of this type of diagnosis.

Techniques for identifying genetic material can obviously be used for the detection of Chlamydia trachomatis. Among these techniques, enzymatic gene amplification or PCR is favoured by those skilled in the art. The technique indeed makes it possible to identify Chlamydia trachomatis with a very high sensitivity and complete specificity. Initially used in specialist laboratories, PCR is now performed in numerous medical laboratories. This diagnostic approach is important because it allows detection of the bacteria even in samples which have been transported under poor conditions.

The treatment of *Chlamydia* urethritis with antibiotics such as tetracycline or quinolones is very effective. The duration of treatment varies between 7 and 14 days. The treatment of pregnant women poses the problem of contraindications to tetracycline.

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Neonatal infections caused by *Chlamydia trachomatis* are explained by the frequency of these bacteria in the cervix. In some studies, 5% to 13% of impairments are observed in the cervix in asymptomatic pregnant women. The neonates risk, in this case, developing an inclusion conjunctivitis. Not only can *Chlamydia trachomatis* be isolated from the children's eyes, but also persistently from the rhinopharynx and also from the rectum. Pneumopathies and otitis media are also found, a result of contamination at childbirth.

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Differential diagnosis of inclusion conjunctivitis in neonates is required with gonococcal ophthalmia; while the duration of incubation is from one to three days in the case of a gonococcal ophthalmia, neonatal inclusion conjunctivitis has an acute beginning with discharge and formation of membranes or even of conjunctival scars.

Treatment consists of oral erythromycin at the dose of 40 to 50 mg per kg of weight, for two to three weeks. In a nonendemic trachoma region, this disease never progresses to chronicity.

Finally, mention should be made of infantile pneumopathy. The syndrome is well defined; it is found in children affected by *Chlamydia trachomatis*. Less than ten children are affected by *Chlamydia trachomatis* pneumopathies per thousand births. The syndrome is, in this case, always found at an early age (less than four months).

Venereal lymphogranulomatosis is an infection which is transmitted through sexual contact and is due to *Chlamydia trachomatis* strains L1, L2 and L3. In humans, a passing primary genital lesion is followed by an often suppurative and multiple regional lymphadenopathy. This disease is a general disease which is accompanied by fever and a rise in the number of white blood cells. If it progresses to chronicity, the disease then becomes complicated with genital elephantiasis, stricture or even fistula of the genital apparatus, of the penis, of the urethra and of the rectum.

In three Chlamydia trachomatis strains L1, L2 and L3 are responsible for venereal lymphogranulomatosis. These Chlamydia strains are more virulent than the strains responsible for trachoma and STD. It is very important to note that venereal lymphogranulomatosis is a systemic disease which affects primarily the lymphatic tissue. Generally transmitted by the sexual route, Chlamydia trachomatis L may also cause contamination through direct contact or even during poor laboratory handling. In spite of these variable modes of transmission, the age for the highest incidence of these diseases corresponds to that for greater sexual activity. Venereal lymphogranulomatosis is still endemic in South America, in Africa and sometimes in Asia. For a long time, the prevalence of venereal lymphogranulomatosis was difficult to establish because of the difficulty of performing diagnosis with certitude. It should also be noted that men are affected more often than women. In low endemic regions, it is difficult to recognize the reservoir of microbes. This situation is explained by the fact that the isolation of the strains causing venereal lymphogranulomatoses from asymptomatic subjects is rarely successful.

Clinical impairment by venereal lymphogranulomatosis manifests itself by the appearance of a small ulcer 3 to 21 days after the exposure of small nonpainful vesicles. In both men and women, the lesion is most often silent. Since this impairment disappears within a few days and causes no functional discomfort and leaves no visible scar, the disease is often recognized late. The venereal lymphogranulomatosis strains may be found in the urethra or the endocervix in patients with inguinal adenopathies; these regions are then considered as the initial site of infection. The

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characteristic feature of the venereal lymphogranulomatosis strains is that from the initial site of infection, *Chlamydia* exhibits a diffusion drained by the lymphatic ducts. The disease is then complicated by a ganglionic impairment of the region draining the site of inoculation. By way of example, anorectal infection causes deep adenopathies. These adneopathies are marked by the appearance of a periadenitis which forms a fluctuating and suppurative ganglionic mass. Fistulae will appear during the decline of the disease. As general signs are present at this stage of the disease, it is often confused with a malignant lymphoma. The other general complications are rarely observed. Clinical examinations have been able to lead biologists to isolate *Chlamydia* from the cerebrospinal fluid or from the blood. It should also be noted that in a number of cases (5%), venereal lymphogranulomatosis is complicated by a chronic oedema: this is genital elephantiasis.

The diagnosis of venereal lymphogranulomatosis requires the isolation of the Chlamydia strains involved in the disease. However, isolation on cell cultures is rarely used, but immunological reactions may be used.

The treatment of venereal lymphogranulomatosis in its initial phase is identical to the treatment of other *Chlamydia* infections. In the chronic phases, antibiotics have little effect on the progress of the disease, but they are however useful in case of superinfection. Although the recommended therapeutic arsenal is identical, it is advisable to prolong the treatment for a period of at least four weeks. In addition to this treatment, reconstructive surgery may be useful in cases of urethral, penile or rectal strictures, as well as for the treatment of fistulae.

In conclusion, a short and effective treatment, without recurrences, and a well-tolerated treatment of *Chlamydia trachomatis* infections therefore remains desirable.

An even greater need up until now relates to a diagnosis which is specific to each of the strains, which is sensitive, which can be carried out conveniently and rapidly, and which allows early detection of the infection.

No vaccine is currently available against *Chlamydia trachomatis*. The role of the immune defense in the physiology and pathology of the disease should probably be understood in order to develop satisfactory vaccines.

More detailed information relating to the biology of these strains, their interactions with their hosts, the associated phenomena of infectivity and those of escaping the immune defenses of the host in particular, and finally their involvement in the development of the these associated pathologies, will allow a better understanding of these mechanisms. In the light of the preceding text which shows in particular the limitations of the means of controlling *Chlamydia trachomatis* infection, it is therefore at present essential, on the one hand, to develop molecular tools, in particular from a better genetic knowledge of *Chlamydia trachomatis*, but also to develop new preventive and therapeutic treatments, new diagnostic methods and new vaccine strategies which are specific,

effective and tolerated. This is precisely the object of the present invention.

The subject of the present invention is the nucleotide sequence having the sequence SEQ ID No. 1 of the Chlamydia trachomatis LGV2 genome. However, the invention is not limited to SEQ ID No. 1, but encompasses genomes and nucleotides encoding polypeptides of strain variants, 5 polymorphisms, allelic variants, and mutants.

Thus, the subject of the present invention encompasses nucleotide sequences characterized in that they are chosen from:

- the nucleotide sequence of SEQ ID No. 1, a nucleotide sequence exhibiting at least 99.9% identity with the sequence SEQ ID No. 1, the nucleotide sequence of the genomic DNA 10 contained within ECACC Deposit No. 98112618, the nucleotide sequence of a clone insert within ECACC Deposit No. 98112617 (these being provisional deposit numbers);
 - b) a nucleotide sequence homologous to the sequence SEQ ID No. 1;
 - c) a polynucleotide sequence that hybridizes to the nucleotide sequence of a) under conditions of high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and $500~\mu g/ml$ denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 10⁶ cpm of ³²P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.
 - (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are

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well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.

- d) a nucleotide sequence complementary to the sequence SEQ ID No. 1 or complementary to a nucleotide sequence as defined in a), b) or c), and a nucleotide sequence of their corresponding RNA;
- e) a nucleotide sequence of a representative fragment of the sequence SEQ ID No. 1, or of a representative fragment of the nucleotide sequence as defined in a), b), c) or d);
 - f) a nucleotide sequence comprising a sequence as defined in a), b), c), d) or e);
- g) a nucleotide sequence capable of being obtained from a nucleotide sequence as defined in a), b), c), d), e) or f); and
- h) a modified nucleotide sequence of a nucleotide sequence as defined in a), b), c), d), e), f) or g).
- Sequence of the genome, or genomic sequence of *Chlamydia trachomatis* is understood to mean the sequence of the chromosome of *Chlamydia trachomatis*, in contrast with the plasmid sequence of *Chlamydia trachomatis*.

Nucleotide sequence, polynucleotide or nucleic acid are understood to mean, according to the present invention, either a double-stranded DNA, a single-stranded DNA or products of transcription of the said DNAs.

It should be understood that the present invention does not relate to the genomic nucleotide sequences of *Chlamydia trachomatis* taken in their natural environment, that is to say in the natural state. They are sequences which may have been isolated, purified or partially purified, by separation methods such as, for example, ion-exchange chromatography, molecular size exclusion chromatography or affinity chromatography, or alternatively fractionation techniques based on solubility in various solvents, or by genetic engineering methods such as amplification, cloning or subcloning, it being possible for the sequences of the invention to be carried by vectors.

The nucleotide sequence SEQ ID No. 1 was obtained by sequencing the *Chlamydia trachomatis LGV2* genome by the method of directed sequencing after fluorescent automated sequencing of the inserts of clones and assembling of these sequences of nucleotide fragments (inserts) by means of softwares (cf. Examples). In spite of the high precision of the sequence SEQ ID No. 1, it is possible that it does not perfectly, 100% represent the nucleotide sequence of the *Chlamydia trachomatis LGV2* genome and that a few rare sequencing errors or uncertainties still remain in the sequence SEQ ID No. 1. In the present invention, the presence of an uncertainty for an amino acid is designated by "Xaa" and that for a nucleotide is designated by "N" in the sequence

listing below. These few rare errors or uncertainties could be easily detected and corrected by persons skilled in the art using the entire chromosome and/or its representative fragments according to the invention and standard amplification, cloning and sequencing methods, it being possible for the sequences obtained to be easily compared, in particular by means of a computer software and using computer-readable media for recording the sequences according to the invention as described, for example, below. After correcting these possible rare errors or uncertainties, the corrected nucleotide sequence obtained would still exhibit at least 99.9% identity with the sequence SEQ ID No. 1. Such rare sequencing uncertainties are not present within the DNA contained within ECACC Deposit No. 98112617 or 98112618 (provisional numbers) and whatever rare sequence uncertainties that exist within SEQ ID No. 1 can routinely be corrected utilizing the DNA of the ECACC Deposits.

Homologous nucleotide sequence for the purposes of the present invention is understood to mean a nucleotide sequence having a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, this percentage being purely statistical and it being possible for the differences between the two nucleotide sequences to be distributed randomly and over their entire length. The said homologous sequences exhibiting a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, may comprise, for example, the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the Chlamydia family, including the species Chlamydia pneumoniae, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the variants of the species Chlamydia trachomatis. In the present invention, the terms family and genus are mutually interchangeable, the terms variant, serotype, strain and subspecies are also mutually interchangeable. These homologous sequences may thus correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in Chlamydia.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

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In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well known in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

(1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;

(2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database;

(3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;

(4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and

(5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990, *Proc. Natl. Acad. Sci. USA 87*:2267-2268).

Nucleotide sequence complementary to a sequence of the invention is understood to mean any DNA whose nucleotides are complementary to those of the sequence of the invention, and whose orientation is reversed (antiparallel sequence).

The present invention further comprises fragments of the sequences of a) through h) above. Representative fragments of the sequences according to the invention will be understood to mean any nucleotide fragment having at least 8 successive nucleotides, preferably at least 12

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successive nucleotides, and still more preferably at least 15 or at least 20 successive nucleotides of the sequence from which it is derived. It is understood that such fragments refer only to portions of SEQ ID No. 1 that are not currently listed in a publicly available database.

Among these representative fragments, those capable of hybridizing under stringent conditions with a nucleotide sequence according to the invention are preferred. Hybridization under stringent conditions means that the temperature and ionic strength conditions are chosen such that they allow hybridization to be maintained between two complementary DNA fragments.

By way of illustration, high stringency conditions for the hybridization step for the purposes of defining the nucleotide fragments described above, are advantageously the following.

The hybridization is carried out at a preferred temperature of 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15 M NaCl and 0.05 M Na citrate. The washing steps may be, for example, the following:

2 x SSC, 0.1% SDS at room temperature followed by three washes with 1 x SSC, 0.1% SDS; 0.5 x SSC, 0.1% SDS; 0.1 x SSC, 0.1% SDS at 68°C for 15 minutes.

Intermediate stringency conditions, using, for example, a temperature of 60°C in the presence of a 5 x SSC buffer, or of low stringency, for example a temperature of 50°C in the presence of a 5 x SSC buffer, respectively require a lower overall complementarity for the hybridization between the two sequences.

The stringent hybridization conditions described above for a polynucleotide of about 300 bases in size will be adapted by persons skilled in the art for larger- or smaller-sized oligonucleotides, according to the teaching of Sambrook et al., 1989.

Among the representative fragments according to the invention, those which can be used as primer or probe in methods which make it possible to obtain homologous sequences or their representative fragments according to the invention, or to reconstitute a genomic fragment found to be incomplete in the sequence SEQ ID No. 1 or carrying an error or an uncertainty, are also preferred, these methods, such as the polymerase chain reaction (PCR), cloning and sequencing of nucleic acid being well known to persons skilled in the art. These homologous nucleotide sequences corresponding to mutations or to inter- or intra-species variations, as well as the complete genomic sequence or one of its representative fragments capable of being reconstituted, of course form part of the invention.

Among the said representative fragments, those which can be used as primer or probe in methods allowing diagnosis of the presence of *Chlamydia trachomatis* or one of its associated microorganisms as defined below are also preferred.

The representative fragments capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia trachomatis* or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia trachomatis* or one of its associated

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microorganisms in the host cell and/or organism, are also preferred. Replication cycle is intended to designate invasion, multiplication, intracellular localization, in particular retention in the vacuole and inhibition of the process of fusion to the lysosome, and propagation of *Chlamydia trachomatis* or one of its associated microorganisms from host cells to host cells.

Among the said representative fragments, those corresponding to nucleotide sequences corresponding to open reading frames, called ORF sequences (ORF for open reading frame), and encoding polypeptides, such as for example, but without being limited thereto, the ORF sequences which will be later described, are finally preferred.

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The representative fragments according to the invention may be obtained, for example, by specific amplification, such as PCR, or after digestion, with appropriate restriction enzymes, of nucleotide sequences according to the invention; these methods are in particular described in the manual by Sambrook et al., 1989. The said representative fragments may also be obtained by chemical synthesis when they are not too large in size and according to methods well known to persons skilled in the art. For example, such fragments can be obtained by isolating fragments of the genomic DNA of ECACC Deposit No. 98112618 or a clone insert present at this ECACC Deposit No. 98112617 (provisional numbers).

The representative fragments according to the invention may be used, for example, as primer, to reconstitute some of the said representative fragments, in particular those in which a portion of the sequence is likely to be missing or imperfect, by methods well known to persons skilled in the art such as amplification, cloning or sequencing techniques.

Modified nucleotide sequence will be understood to mean any nucleotide sequence obtained by mutagenesis according to techniques well known to persons skilled in the art, and exhibiting modifications in relation to the normal sequences, for example mutations in the regulatory and/or promoter sequences for the expression of a polypeptide, in particular leading to a modification of the level of expression of the said polypeptide or to a modulation of the replicative cycle.

Modified nucleotide sequence will also be understood to mean any nucleotide sequence encoding a modified polypeptide as defined below.

The subject of the present invention also includes *Chlamydia trachomatis* nucleotide sequences characterized in that they are chosen from a nucleotide sequence of an open reading frame (ORF), that is, the ORF2 to ORF1197 sequences.

The ORF2 to ORF1197 nucleotide sequences are defined in Tables 1 and 2, infra, represented below by their position on the sequence SEQ ID No. 1. For example, the ORF10 sequence is defined by the nucleotide sequence between the nucleotides at position 9828 and 10430 on the sequence SEQ ID No. 1, ends included. ORF2 to ORF1197 have been identified via homology analyses as well as via analyses of potential ORF start sites, as discussed in the examples

below. It is to be understood that each identified ORF of the invention comprises a nucleotide sequence that spans the contiguous nucleotide sequence from the codon immediately 3' to the stop codon of the preceding ORF and through the 5' codon to the next stop codon of SEQ ID No.:1 inframe to the ORF nucleotide sequence. Table 2, *infra*, lists the beginning, end and potential start site of each of ORFs 2-1197. In one embodiment, the ORF comprises the contiguous nucleotide sequence spanning from the potential ORF start site downstream (that is, 3') to the ORF stop codon (or the ORF codon immediately adjacent to and upstream of the ORF stop codon). ORF2 to ORF1197 encode the polypeptides of SEQ ID No. 2 to SEQ ID No. 1197.

Upon introduction of minor frameshifts, certain individual ORFs can comprise larger

«combined» ORFs. A list of such putative «combined» ORFs is shown in Table 3, below. For example, a combined ORF can comprise ORF 1076 and ORF 1073, including intervening in-frame, nucleotide sequences. The order of ORFs (5' to 3'), within each «combined» ORF is as listed. It is to be understood that when ORF2 to ORF1197 are referred to herein, such reference is also meant to include «combined» ORFs. Polypeptide sequences encoded by such «combined» ORFs are also part of the present invention.

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Table 3
    ORF 1076, ORF 1073;
    ORF 3, ORF 2:
    ORF 23, ORF 22, ORF 21;
20 ORF 1141, ORF 477, ORF 478, ORF 479;
    ORF 487, ORF 486, ORF 485, ORF 484, ORF 483, ORF 482, ORF 481;
    ORF 488, ORF 489;
    ORF 573, ORF 572, ORF 571;
    ORF 817, ORF 818;
25 ORF 819, ORF 820;
    ORF 1037, ORF 1038:
    ORF 1071, ORF 1070;
    ORF 17, ORF 1077;
    ORF 1185, ORF 933, ORF 934;
30 ORF 1060, ORF 1059;
    ORF 155, ORF 156;
    ORF 679, ORF 680;
    ORF 879, ORF 878;
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ORF 1028; ORF 1029, 35 and representative fragments. Table 1 also depicts the results of homology searches that compared the sequences of the polypeptides encoded by each of the ORFs to sequences present in public published databases. It is understood that in one embodiment, those polypeptides listed in Table 1 as exhibiting greater than about 95% identity to a polypeptide present in a publicly disclosed database are not considered part of the present invention; likewise in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention. In another embodiment, it is understood that those polypeptides listed in Table 1 as exhibiting greater than about 99% identity to a polypeptide present in a publicly disclosed database are not considered part of the invention; likewise, in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention.

The invention also relates to the nucleotide sequences characterized in that they comprise a nucleotide sequence chosen from:

- a) an ORF2 to ORF1197, a «combined» ORF nucleotide sequence, the nucleotide sequence of the genomic DNA contained within ECACC Deposit No. 98112618 or the nucleotide sequence of a clone insert in ECACC Deposit No. 98112617 according to the invention;
- b) a homologous nucleotide sequence exhibiting at least 80% identity across an entire ORF2 to ORF1197 nucleotide sequence according to the invention or as defined in a);
- c) a polynucleotide sequence that hybridizes to ORF2 to ORF1197 under conditions of high or intermediate stringency as described below:
- 20 (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and $500~\mu\text{g/ml}$ denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 μg/ml denatured salmon sperm DNA and 5-20 X 106 cpm of ³²P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.

Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a *Chlamydia trachomatis* polypeptide.

- (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.
- d) a complementary or RNA nucleotide sequence corresponding to an ORF2 to ORF1197 sequence according to the invention or as defined in a), b) or c);
 - e) a nucleotide sequence of a representative fragment of an ORF2 to ORF1197 sequence according to the invention or of a sequence as defined in a), b), c) or d);
 - f) a nucleotide sequence capable of being obtained from an ORF2 to ORF1197 sequence according to the invention or as defined in a), b), c), d) or e); and
- 20 g) a modified nucleotide sequence of an ORF2 to ORF1197 sequence according to the invention or as defined in a), b), c), d), e) or f).

As regards the homology with the ORF2 to ORF1197 nucleotide sequences, the homologous sequences exhibiting a percentage identity with the bases of one of the ORF2 to ORF1197 nucleotide sequences of at least 80%, preferably 90% and 95%, are preferred. Such homologous sequences are identified routinely via, for example, the algorithms described above and in the examples below. The said homologous sequences correspond to the homologous sequences as defined above and may comprise, for example, the sequences corresponding to the ORF sequences of a bacterium belonging to the Chlamydia family, including the species Chlamydia pneumoniae, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the ORF sequences of a bacterium belonging to the variants of the species Chlamydia trachomatis. These homologous sequences may likewise correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in

Chlamydia.

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The invention comprises the polypeptides encoded by a nucleotide sequence according to the invention, preferably by a representative fragment of the sequence SEQ ID No. 1 and corresponding to an ORF sequence, in particular the Chlamydia trachomatis polypeptides, 5 characterized in that they are chosen from the sequences SEQ ID No. 2 to SEQ ID No. 1197, and representative fragments thereof. However, the invention is not limited to polypeptides encoded by ORFs in SEQ ID No. 1 and its corresponding ORF sequences, but encompasses polypeptides of strain variants, polymorphisms, allelic variants, and mutants.

Thus, the invention also comprises the polypeptides characterized in that they 10 comprise a polypeptide chosen from:

- a polypeptide encoded by a polynucleotide sequence in SEQ ID No. 1 (e.g., any a) polypeptide encoded by a polynucleotide sequence corresponding to ORF2 to ORF1197) and/or representative fragments thereof according to the invention;
- a polypeptide homologous to a polypeptide according to the invention, or as defined b) 15 in a);
 - c) a polypeptide encoded by a polynucleotide sequence that hybridizes to SEQ ID No. 1 or ORF2 to ORF1197 under high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 106 cpm of 32P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 25 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other 30 conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197 . In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.

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- (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.
 - d) a fragment of at least 5 amino acids of a polypeptide according to the invention, or as defined in a), b) or c);
- e) a biologically active fragment of a polypeptide according to the invention, or as defined in a), b), c) or d); and
 - f) a modified polypeptide of a polypeptide according to the invention, as defined in a), b), c), d) or e).

In the present description, the terms polypeptide, peptide and protein are interchangeable.

It should be understood that the invention does not relate to the polypeptides in natural form, that is to say that they are not taken in their natural environment but that they may have been isolated or obtained by purification from natural sources, or alternatively obtained by genetic recombination, or else by chemical synthesis and that they may, in this case, comprise nonnatural amino acids, as will be described below.

Homologous polypeptide will be understood to designate the polypeptides exhibiting, in relation to the natural polypeptide, certain modifications such as in particular a deletion, addition or substitution of at least one amino acid, a truncation, an extension, a chimeric fusion, and/or a mutation, or polypeptides exhibiting post-translational modifications. Among the homologous polypeptides, those whose amino acid sequence exhibits at least 80%, preferably 90%, homology or identity with the amino acid sequences of the polypeptides according to the invention are preferred. In the case of a substitution, one or more consecutive or nonconsecutive amino acids are replaced by "equivalent" amino acids. The expression "equivalent" amino acid is intended here to designate any amino acid capable of being substituted for one of the amino acids in the basic structure without, however, essentially modifying the biological activities of the corresponding peptides and as will be defined later.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well know in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

- (1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence 15 database;
 - (2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database;
 - (3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
 - (5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure,

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g.,

Washington: National Biomedical Research Foundation)

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Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268).

Equivalent amino acids may be determined either based on their structural homology with the amino acids for which they are substituted, or on results of comparative tests of biological activity between the various polypeptides which may be carried out.

By way of example, there may be mentioned the possibilities of substitutions which may be carried out without resulting in a substantial modification of the biological activity of the corresponding modified polypeptides; the replacements, for example, of leucine with valine or isoleucine, of aspartic acid with glutamic acid, of glutamine with asparagine, of arginine with lysine, and the like, the reverse substitutions naturally being feasible under the same conditions.

The homologous polypeptides also correspond to the polypeptides encoded by the homologous nucleotide sequences as defined above and thus comprise in the present definition the mutated polypeptides or polypeptides corresponding to inter- or intra-species variations which may exist in *Chlamydia*, and which correspond in particular to truncations, substitutions, deletions and/or additions of at least one amino acid residue.

Biologically active fragment of a polypeptide according to the invention will be understood to designate in particular a polypeptide fragment, as defined below, exhibiting at least one of the characteristics of the polypeptides according to the invention, in particular in that it is:

- capable of eliciting an immune response directed against Chlamydia trachomatis; and/or
- capable of being recognized by an antibody specific for a polypeptide according to the 20 invention; and/or
 - capable of binding to a polypeptide or to a nucleotide sequence of *Chlamydia trachomatis*; and/or
- capable of modulating, regulating, inducing or inhibiting the expression of a gene of Chlamydia trachomatis or one of its associated microorganisms, and/or capable of modulating the
 replication cycle of Chlamydia trachomatis or one of its associated microorganisms in the host cell and/or organism; and/or
 - capable of generally exerting an even partial physiological activity, such as for example a structural activity (cellular envelope, ribosome), an enzymatic (metabolic) activity, a transport activity, an activity in the secretion or in the virulence.

A representative polypeptide fragment according to the invention is understood to designate a polypeptide comprising a minimum of 5 amino acids, preferably 10 amino acids or preferably 15 amino acids. It is to be understood that such fragments refer only to portions of polypeptides encoded by ORF2 or ORF1197 that are not currently listed in a publicly available database.

The polypeptide fragments according to the invention may correspond to isolated or

purified fragments which are naturally present in Chlamydia trachomatis or which are secreted by Chlamydia trachomatis, or may correspond to fragments capable of being obtained by cleaving the said polypeptide with a proteolytic enzyme, such as trypsin or chymotrypsin or collagenase, or with a chemical reagent, such as cyanogen bromide (CNBr) or alternatively by placing the said polypeptide in a highly acidic environment, for example at pH 2.5. Such polypeptide fragments may be equally well prepared by chemical synthesis, using hosts transformed with an expression vector according to the invention containing a nucleic acid allowing the expression of the said fragments, placed under the control of appropriate elements for regulation and/or expression.

«Modified polypeptide» of a polypeptide according to the invention is understood to designate a polypeptide obtained by genetic recombination or by chemical synthesis as will be described below, exhibiting at least one modification in relation to the normal sequence. These modifications may in particular affect amino acids responsible for a specificity or for the efficiency of the activity, or responsible for the structural conformation, for the charge or for the hydrophobicity, and for the capacity for multimerization and for membrane insertion of the polypeptide according to the invention. It is thus possible to create polypeptides with an equivalent, an increased or a reduced activity, and with an equivalent, a narrower or a broader specificity. Among the modified polypeptides, there may be mentioned the polypeptides in which up to 5 amino acids may be modified, truncated at the N- or C-terminal end, or alternatively deleted, or else added.

As is indicated, the modifications of the polypeptide may have in particular the 20 objective:

- of making it capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia*, in particular of *Chlamydia trachomatis* and its variants, or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia*, in particular of *Chlamydia trachomatis* and its variants, or one of its associated microorganisms, in the host cell and/or organism,
- of allowing its use in methods of biosynthesis or of biodegradation, or its incorporation into vaccine compositions,
- of modifying its bioavailability as a compound for therapeutic use.

The said modified polypeptides may also be used on any cell or microorganism for which the said modified polypeptides will be capable of modulating, regulating, inhibiting or inducing gene expression, or of modulating the growth or the replication cycle of the said cell or of the said microorganism. The methods allowing demonstration of the said modulations on eukaryotic or prokaryotic cells are well known to persons skilled in the art. The said cells or microorganisms will be chosen, in particular, from tumour cells or infectious microorganisms and the said modified polypeptides may be used for the prevention or treatment of pathologies linked to the presence of the

said cells or of the said microorganisms. It is also clearly understood that the nucleotide sequences encoding the said modified polypeptides may be used for the said modulations, for example by the intermediacy of vectors according to the invention and which are described below, so as to prevent or to treat the said pathologies.

The above modified polypeptides may be obtained using combinatory chemistry, in which it is possible to systematically vary portions of the polypeptide before testing them on models, cell cultures or microorganisms for example, so as to select the compounds which are the most active or which exhibit the desired properties.

Chemical synthesis also has the advantage of being able to use:

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- nonnatural amino acids, or
- nonpeptide bonds.

Accordingly, in order to extend the life of the polypeptides according to the invention, it may be advantageous to use nonnatural amino acids, for example in the D form, or alternatively amino acid analogues, in particular sulphur-containing forms for example.

Finally, the structure of the polypeptides according to the invention, its homologous or modified forms, as well as the corresponding fragments may be integrated into chemical structures of the polypeptide type and the like. Accordingly, it may be advantageous to provide at the N- and C-terminal ends compounds which are not recognized by proteases.

Also forming part of the invention are the nucleotide sequences encoding a polypeptide according to the invention. Described below are ORF nucleotide sequences encoding polypeptides exhibiting particularly preferable characteristics. For each group of preferred ORFs described below, it is to be understood that in addition to the individual ORFs listed, in instances wherein such ORFs are present as part of «combined» ORFs, the «combined» ORFs are also to be included within the preferred group.

More particularly, the subject of the invention is nucleotide sequences, characterized in that they encode a polypeptide of the cellular envelope, preferably of the outer cellular envelope of Chlamydia trachomatis or one of its representative fragments, such as for example the predominant proteins of the outer membrane, the adhesion proteins or the proteins entering into the composition of the Chlamydia wall. Among these sequences, the sequences comprising a nucleotide sequence chosen 30 from the following sequences are most preferred:

ORF3; ORF19; ORF51; ORF189; ORF212; ORF213; ORF324; ORF477; ORF478; ORF479; ORF481; ORF482; ORF483; ORF484; ORF486; ORF488; ORF489; ORF490; ORF572; ORF573; ORF742; ORF817; ORF818; ORF820; ORF1035; ORF1036; ORF1037; ORF1038; ORF1070; ORF1071; ORF1073 and one of their representative fragments.

35 The structure of the cytoplasmic membranes and of the wall of bacteria is dependent on the associated proteins. The structure of the cytoplasmic membrane makes it impermeable to water, to water-soluble substances and to small-sized molecules (ions, small inorganic molecules, peptides or proteins). To enter into or to interfere with a cell or a bacterium, a ligand must establish a special relationship with a protein anchored in the cytoplasmic membrane (the receptor). These proteins which are anchored on the membrane play an important role in metabolism since they control the exchanges in the bacterium. These exchanges apply to molecules of interest for the bacterium (small molecules such as sugars and small peptides) as well as undesirable molecules for the bacterium such as antibiotics or heavy metals.

The double lipid layer structure of the membrane requires the proteins which are inserted therein to have hydrophobic domains of about twenty amino acids forming an alpha helix. Predominantly hydrophobic and potentially transmembrane regions may be predicted from the primary sequence of the proteins, itself deduced from the nucleotide sequence. The presence of one or more putative transmembrane domains raises the possibility for a protein to be associated with the cytoplasmic membrane and to be able to play an important metabolic role therein or alternatively for the protein thus exposed to be able to exhibit potentially protective epitopes.

If the proteins inserted into the membrane exhibit several transmembrane domains capable of interacting with one another via electrostatic bonds, it then becomes possible for these proteins to form pores which go across the membrane which becomes permeable for a number of substances. It should be noted that proteins which do not have transmembrane domains may also be anchored by the intermediacy of fatty acids in the cytoplasmic membrane, it being possible for the breaking of the bond between the protein and its anchor in some cases to be responsible for the release of the peptide outside the bacterium.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF2; ORF3; ORF5; ORF8; ORF9; ORF10; ORF11; ORF12; ORF17; ORF21; ORF26; ORF27; ORF28; ORF29; ORF30; ORF31; ORF33; ORF35; ORF37; ORF39; ORF40; ORF41; ORF42; ORF43; ORF44; ORF45; ORF46; ORF47; ORF48; ORF49; ORF52; ORF53; ORF55; ORF56; ORF58; ORF66; ORF68; ORF70; ORF74; ORF75; ORF76; ORF78; ORF79; ORF81; ORF82; ORF83; ORF86; ORF91; ORF92; ORF94; ORF97; ORF100; ORF102; ORF103; ORF105; ORF106; ORF107; ORF109; ORF110; ORF111; ORF112; ORF113; ORF114; ORF115; ORF116; ORF117; ORF120; ORF122; ORF123; ORF130; ORF134; ORF135; ORF137; ORF140; ORF141; ORF143; ORF144; ORF145; ORF147; ORF148; ORF149; ORF150; ORF151; ORF155; ORF156; ORF162; ORF163; ORF164; ORF165; ORF166; ORF167; ORF168; ORF169; ORF170; ORF171;

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ORF173; ORF175; ORF176; ORF187; ORF181; ORF183; ORF184; ORF186; ORF187; ORF188;
    ORF190; ORF191; ORF192; ORF194; ORF195; ORF196; ORF197; ORF198; ORF199; ORF201;
    ORF202; ORF204; ORF206; ORF207; ORF209; ORF212; ORF213; ORF217; ORF219; ORF220;
    ORF221; ORF222; ORF223; ORF224; ORF225; ORF227; ORF228; ORF231; ORF232; ORF234;
 5 ORF236; ORF243; ORF244; ORF245; ORF247; ORF248; ORF249; ORF252; ORF254;
    ORF257; ORF260; ORF261; ORF263; ORF265; ORF266; ORF267; ORF270; ORF271; ORF272;
    ORF274; ORF276; ORF277; ORF278; ORF282; ORF283; ORF284; ORF285; ORF287;
    ORF289; ORF290; ORF291; ORF294; ORF298; ORF305; ORF306; ORF310; ORF311; ORF313;
    ORF315; ORF316; ORF319; ORF320; ORF322; ORF323; ORF325; ORF326; ORF327; ORF328;
10 ORF330; ORF331; ORF332; ORF333; ORF334; ORF335; ORF336; ORF338; ORF339; ORF340;
    ORF341; ORF344; ORF345; ORF348; ORF349; ORF350; ORF351; ORF352; ORF353; ORF356;
    ORF357; ORF358; ORF361; ORF362; ORF366; ORF367; ORF368; ORF370; ORF372; ORF373;
    ORF375; ORF377; ORF378; ORF380; ORF382; ORF383; ORF384; ORF385; ORF387;
    ORF389; ORF390; ORF391; ORF393; ORF396; ORF398; ORF399; ORF403; ORF404; ORF406;
15 ORF407; ORF413; ORF414; ORF417; ORF418; ORF420; ORF421; ORF424; ORF426; ORF427;
    ORF428; ORF430; ORF433; ORF434; ORF435; ORF436; ORF437; ORF440; ORF443; ORF446;
    ORF448; ORF450; ORF451; ORF454; ORF455; ORF457; ORF458; ORF459; ORF463; ORF464;
    ORF466; ORF467; ORF468; ORF469; ORF470; ORF473; ORF474; ORF475; ORF476; ORF477;
    ORF479; ORF480; ORF481; ORF483; ORF484; ORF485; ORF486; ORF487; ORF488; ORF491;
   ORF493; ORF496; ORF497; ORF498; ORF500; ORF501; ORF503; ORF504; ORF508; ORF512;
    ORF513; ORF514; ORF519; ORF521; ORF523; ORF524; ORF526; ORF527; ORF529; ORF530;
    ORF531; ORF532; ORF534; ORF536; ORF537; ORF538; ORF540; ORF541; ORF542; ORF543;
    ORF544; ORF545; ORF546; ORF5547; ORF551; ORF5552; ORF5553; ORF5555; ORF5558; ORF5559;
    ORF560; ORF561; ORF562; ORF566; ORF567; ORF568; ORF569; ORF571; ORF572; ORF574;
25 ORF575; ORF580; ORF582; ORF585; ORF587; ORF589; ORF592; ORF593; ORF595;
    ORF596; ORF597; ORF599; ORF601; ORF602; ORF603; ORF604; ORF608; ORF609; ORF610;
    ORF611; ORF615; ORF616; ORF617; ORF618; ORF621; ORF622; ORF623; ORF624; ORF625;
    ORF628; ORF632; ORF633; ORF634; ORF635; ORF637; ORF638; ORF640; ORF641; ORF643;
    ORF646; ORF648; ORF651; ORF652; ORF653; ORF655; ORF658; ORF664;
30 ORF665; ORF666; ORF669; ORF670; ORF671; ORF672; ORF673; ORF674; ORF676;
    ORF677; ORF678; ORF680; ORF682; ORF683; ORF684; ORF686; ORF688; ORF689; ORF690;
    ORF691; ORF692; ORF693; ORF695; ORF696; ORF698; ORF701; ORF703; ORF704; ORF705;
   ORF706; ORF707; ORF709; ORF710; ORF711; ORF712; ORF713; ORF714; ORF715; ORF717;
   ORF718; ORF720; ORF721; ORF722; ORF724; ORF726; ORF728; ORF729; ORF730; ORF731;
   ORF732; ORF733; ORF734; ORF737; ORF738; ORF739; ORF740; ORF742; ORF744;
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ORF745; ORF746; ORF750; ORF751; ORF752; ORF753; ORF754; ORF755; ORF757; ORF758; ORF759; ORF760; ORF764; ORF766; ORF768; ORF769; ORF771; ORF772; ORF773; ORF774; ORF775; ORF776; ORF777; ORF778; ORF779; ORF780; ORF781; ORF782; ORF783; ORF786; ORF787; ORF788; ORF789; ORF790; ORF793; ORF798; ORF800; ORF802; ORF803; 5 ORF806; ORF809; ORF810; ORF811; ORF813; ORF814; ORF817; ORF820; ORF822; ORF824; ORF825; ORF827; ORF828; ORF829; ORF830; ORF833; ORF834; ORF835; ORF837; ORF838; ORF839; ORF840; ORF841; ORF842; ORF843; ORF845; ORF848; ORF849; ORF850; ORF851; ORF852; ORF855; ORF856; ORF857; ORF859; ORF860; ORF862; ORF863; ORF864; ORF866; ORF869; ORF872; ORF873; ORF874; ORF878; ORF879; ORF880; ORF881; 10 ORF883; ORF884; ORF885; ORF886; ORF887; ORF892; ORF893; ORF894; ORF895; ORF897; ORF899; ORF900; ORF901; ORF904; ORF906; ORF909; ORF910; ORF912; ORF914; ORF917; ORF920; ORF921; ORF922; ORF923; ORF924; ORF925; ORF926; ORF927; ORF930; ORF933; ORF934; ORF935; ORF936; ORF937; ORF940; ORF941; ORF942; ORF943; ORF944; ORF945; ORF947; ORF948; ORF951; ORF952; ORF953; ORF954; ORF955; ORF956; ORF957; ORF958; ORF960; ORF961; ORF962; ORF963; ORF964; ORF966; ORF967; ORF969; ORF970; ORF971; ORF973; ORF974; ORF980; ORF981; ORF982; ORF984; ORF988; ORF989; ORF990; ORF991; ORF995; ORF996; ORF999; ORF1001; ORF1003; ORF1004; ORF1005; ORF1006; ORF1007; ORF1009; ORF1010; ORF1011; ORF1012; ORF1013; ORF1014; ORF1016; ORF1017; ORF1018; ORF1020; ORF1021; ORF1025; ORF1026; ORF1027; ORF1029; ORF1030; ORF1031; ORF1035; ORF1036; ORF1037; ORF1038; ORF1039; ORF1040; ORF1044; ORF1045; ORF1047; ORF1048; ORF1050; ORF1051; ORF1052; ORF1053; ORF1055; ORF1056; ORF1057; ORF1058; ORF1061; ORF1062; ORF1063; ORF1064; ORF1065; ORF1066; ORF1068; ORF1069; ORF1072; ORF1074; ORF1076 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 4 and 6 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF7; ORF14; ORF16; ORF32; ORF34; ORF36; ORF38; ORF50; ORF57; ORF59; ORF61; ORF62; ORF63; ORF64; ORF67; ORF69; ORF72; ORF77; ORF80; ORF84; ORF87; ORF93; ORF95; ORF99; ORF108; ORF119; ORF125; ORF126; ORF129; ORF131; ORF136; ORF139; ORF146; ORF152; ORF154; ORF160; ORF161; ORF172; ORF179; ORF182; ORF185; ORF200; ORF203; ORF205; ORF239; ORF242; ORF250; ORF253; ORF256; ORF259; ORF262; ORF268; ORF275; ORF281; ORF286; ORF288; ORF292; ORF295; ORF296; ORF297; ORF299; ORF300; ORF308; ORF314; ORF317; ORF318; ORF324; ORF342; ORF343; ORF355; ORF360; ORF374; ORF376; ORF386; ORF388; ORF392; ORF394; ORF395; ORF402; ORF405; ORF411; ORF415; ORF416;

ORF422; ORF423; ORF429; ORF432; ORF441; ORF442; ORF444; ORF449; ORF452; ORF456; ORF460; ORF461; ORF465; ORF471; ORF472; ORF482; ORF489; ORF492; ORF494; ORF495; ORF502; ORF505; ORF506; ORF509; ORF516; ORF517; ORF520; ORF525; ORF533; ORF539; ORF549; ORF554; ORF557; ORF563; ORF570; ORF573; ORF581; ORF590; ORF591; ORF600; ORF607; ORF612; ORF613; ORF620; ORF626; ORF629; ORF630; ORF639; ORF644; ORF647; ORF656; ORF659; ORF661; ORF685; ORF687; ORF699; ORF700; ORF708; ORF716; ORF719; ORF725; ORF747; ORF749; ORF756; ORF765; ORF767; ORF794; ORF796; ORF797; ORF799; ORF801; ORF807; ORF821; ORF823; ORF826; ORF847; ORF853; ORF861; ORF870; ORF871; ORF875; ORF882; ORF888; ORF889; ORF902; ORF903; ORF911; ORF916; ORF931; ORF939; ORF975; ORF976; ORF978; ORF983; ORF986; ORF987; ORF992; ORF993; ORF1000; ORF1002; ORF1008; ORF1019; ORF1022; ORF1032; ORF1034; ORF1046; ORF1054; ORF1060; ORF1071 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF4; ORF6; ORF13; ORF20; ORF51; ORF71; ORF88; ORF118; ORF128; ORF132; ORF133; ORF158; ORF159; ORF174; ORF180; ORF189; ORF210; ORF211; ORF214; ORF215; ORF226; ORF229; ORF233; ORF235; ORF240; ORF246; ORF251; ORF255; ORF273; ORF354; ORF364; ORF369; ORF371; ORF397; ORF401; ORF409; ORF412; ORF419; ORF439; ORF453; ORF462; ORF490; ORF510; ORF511; ORF518; ORF535; ORF548; ORF550; ORF564; ORF565; ORF578; ORF579; ORF614; ORF631; ORF636; ORF650; ORF662; ORF667; ORF679; ORF681; ORF702; ORF727; ORF741; ORF763; ORF791; ORF792; ORF815; ORF816; ORF832; ORF846; ORF858; ORF865; ORF867; ORF868; ORF877; ORF891; ORF896; ORF907; ORF908; ORF918; ORF919; ORF932; ORF959; ORF977; ORF994; ORF998; ORF1024; ORF1028; ORF1042; ORF1067; ORF1070; ORF1073 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* surface exposed polypeptide (e.g., an outer membrane protein) or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 2, ORF 3, ORF 21, ORF 22, ORF 23, ORF 53, ORF 77, ORF 187, ORF 203, ORF 383, ORF 477, ORF 478, ORF 479, ORF 481, ORF 482, ORF 483, ORF 484, ORF 485, ORF 486, ORF 487, ORF 488, ORF 489, ORF 490, ORF 571, ORF 572, ORF 573, ORF 593, ORF 670, ORF 693, ORF 742, ORF 749, ORF 801, ORF 817, ORF 818, ORF 819, ORF 820, ORF 851, ORF 902, ORF 923,

35 ORF 1035, ORF 1036, ORF 1037, ORF 1038, ORF 1069, ORF 1070, ORF 1071, ORF 1073, ORF

1076, ORF 1095, ORF 1096, ORF 1141, ORF 1181, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* lipoprotein or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 29, ORF 42, ORF 66, ORF 72, ORF 76, ORF 78, ORF 148, ORF 154, ORF 180, ORF 182, ORF 184, ORF 187, ORF 200, ORF 242, ORF 245, ORF 250, ORF 253, ORF 272, ORF 274, ORF 275, ORF 308, ORF 350, ORF 362, ORF 383, ORF 394, ORF 396, ORF 399,

ORF 422, ORF 488, ORF 535, ORF 568, ORF 573, ORF 578, ORF 593, ORF 607, ORF 625, ORF 662, ORF 669, ORF 688, ORF 690, ORF 716, ORF 773, ORF 778, ORF 781, ORF 783, ORF 788, ORF 817, ORF 848, ORF 851, ORF 853, ORF 857, ORF 875, ORF 877, ORF 886, ORF 898, ORF 902, ORF 923, ORF 938, ORF 976, ORF 978, ORF 990, ORF 1005, ORF 1021, ORF 1035, ORF 1069, ORF 1083, ORF 1088, ORF 1089, ORF 1091, ORF 1092, ORF 1095, ORF 1096, ORF 1100, ORF 1105, ORF 1108, ORF 1117, ORF 1120, ORF 1121, ORF 1124, ORF 1128, ORF 1133, ORF 1135, ORF 1139, ORF 1140, ORF 1157, ORF 1159, ORF 1163, ORF 1165, ORF 1167, ORF 1168, ORF 1169, ORF 1171, ORF 1173, ORF 1174, ORF 1177, ORF 1180, ORF 1181, ORF 1186, ORF 1194, ORF 1197, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide involved in lipopolysaccharide (LPS) biosynthesis, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 17, ORF 201, ORF 691, ORF 807, ORF 936, ORF 983, ORF 1019, ORF 1077 and one of their representative fragments.

Preferably the invention relates to additional LPS-related nucleotide sequences according to the invention, characterized in that they encode:

- 25 (a) a Chlamydia trachomatis KDO (3-deoxy-D-manno-octulosonic acid)-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 41, ORF 242, ORF 269, ORF 772, and one of their representative fragments;
 - (b) a Chlamydia trachomatis phosphomannomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequence: ORF 139, and one of its representative fragments;
 - (c) a Chlamydia trachomatis phosphoglucomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequence: ORF 567, and one of its representative fragments; and
 - (d) a Chlamydia trachomatis lipid A component-related polypeptide or one of its

representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 4, ORF 933, ORF 934, ORF 935, ORF 1185, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* Type III or other, non-Type III secreted polypeptides or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 180, ORF 181, ORF 207, ORF 208, ORF 372, ORF 391, ORF 399, ORF 477, ORF 486, ORF 749, ORF 758, ORF 819, ORF 878, ORF 888, ORF 896, ORF 897, ORF 900, ORF 902, ORF 923, ORF 1015, ORF 1018, ORF 1059, ORF 1060, ORF 1069, ORF 1071, ORF 1073, ORF 1076, ORF 1189, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide containing RGD (Arg-Gly-Asp) attachment sites or one of its representative fragments:

- 15 (a) RGD-containing proteins that are outer membrane proteins, are more likely to play a role in cell attachment. ORFs that encoded a protein containing an RGD sequence and also were classified as outer membrane proteins are ORF 488, ORF 489, ORF 571, ORF 572, ORF 573 or ORF 716, and its representative fragments.
- 20 (b) The outer membrane of Chlamydia is made of cysteine-rich proteins that form a network of both intra and inter molecular disulfide links. This contributes to the integrity of the membrane since Chlamydia lacks the peptidoglycan layer that other gram-negative bacteria have. Cysteine-rich proteins that have the RGD sequence are also considered to be potential vaccine candidates. Cysteine-rich proteins were defined as proteins that had more than 3.0% cysteine in their primary amino acid sequence, above the mean genomic ORF cysteine content. The corresponding ORF is: ORF 1144 and one of its representative fragments.
- (c) The outer membrane of Chlamydia may also contain small proteins that have cysteines in their N- and C-terminus that may contribute to the network formed by disulfide linkages.

 These proteins may be anchored in the outer membrane via their N-terminus and may have their C-terminus exposed, which then can interact with the host cells. Alternatively, these proteins may be anchored in the outer membrane via both N-and C-terminus and may have regions in the middle that may be exposed which can in turn interact with the host cells. ORFs encoding polypeptides that contain cysteines in their first 30 amino acids and also contain an RGD sequence are: ORF 101, ORF 122, ORF 308, ORF 488, ORF 489, ORF 571, ORF 572, ORF 573, ORF 651, ORF 679, ORF 680,

ORF 705, ORF 716, ORF 763, ORF 870, ORF 878, ORF 879, ORF 995, ORF 1028, ORF 1029, ORF 1176, and one of their representative fragments.

(d) RGD-containing ORFs homologous to RGD-containing ORFs from 5 Chlamydia pneumoniae are:
ORF 28, ORF 101, ORF 125, ORF 155, ORF 156, ORF 286, ORF 571, ORF 572, ORF 573, ORF

763, ORF 870, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* cell wall anchored surface polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 662, ORF 681, ORF 1182, ORF 1192, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode Chlamydia trachomatis polypeptides not found in Chlamydia pneumoniae (Blastp P>e⁻¹⁰), said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 2, ORF 18, ORF 60, ORF 66, ORF 67, ORF 68, ORF 69, ORF 70, ORF 81, ORF 89, ORF 107, ORF 108, ORF 109, ORF 134, ORF 147, ORF 191, ORF 194, ORF 216, ORF 217, ORF 218, ORF 219, ORF 220, ORF 221, ORF 222, ORF 223, ORF 224, ORF 225, ORF 228, ORF 235, ORF 257, ORF 276, ORF 277, ORF 278, ORF 279, ORF 280, ORF 281, ORF 282, ORF 283, ORF 284, ORF 285, ORF 289, ORF 291, ORF 298, ORF 313, ORF 314, ORF 315, ORF 316, ORF 334, ORF 335, ORF 336, ORF 337, ORF 338, ORF 339, ORF 340, ORF 381, ORF 393, ORF 413, ORF 418, ORF 419, ORF 420, ORF 421, ORF 422, ORF 423, ORF 436, ORF 460, ORF 475, ORF 476, ORF 480, ORF 485, ORF 487, ORF 491, ORF 492, ORF 493, ORF 494, 25 ORF 496, ORF 500, ORF 504, ORF 514, ORF 527, ORF 559, ORF 569, ORF 570, ORF 575, ORF 580, ORF 582, ORF 593, ORF 598, ORF 632, ORF 640, ORF 651, ORF 671, ORF 690, ORF 694, ORF 698, ORF 710, ORF 722, ORF 723, ORF 724, ORF 770, ORF 771, ORF 782, ORF 783, ORF 784, ORF 790, ORF 795, ORF 798, ORF 805, ORF 810, ORF 817, ORF 829, ORF 830, ORF 864, ORF 866, ORF 876, ORF 887, ORF 892, ORF 899, ORF 913, ORF 921, ORF 933, ORF 938, ORF 30 949, ORF 956, ORF 1010, ORF 1017, ORF 1018, ORF 1027, ORF 1030, ORF 1037, ORF 1038, ORF 1047, ORF 1072, ORF 1074, ORF 1075, ORF 1078, ORF 1079, ORF 1081, ORF 1083, ORF 1084, ORF 1087, ORF 1088, ORF 1089, ORF 1091, ORF 1092, ORF 1094, ORF 1095, ORF 1096, ORF 1098, ORF 1104, ORF 1105, ORF 1106, ORF 1108, ORF 1110, ORF 1114, ORF 1115, ORF 1116, ORF 1117, ORF 1119, ORF 1128, ORF 1132, ORF 1133, ORF 1135, ORF 1136, ORF 1139, ORF 1140, ORF 1141, ORF 1142, ORF 1144, ORF 1148, ORF 1151, ORF 1155, ORF 1157, ORF 1159,

ORF 1161, ORF 1162, ORF 1165, ORF 1166, ORF 1167, ORF 1168, ORF 1169, ORF 1171, ORF 1172, ORF 1173, ORF 1174, ORF 1175, ORF 1176, ORF 1177, ORF 1178, ORF 1180, ORF 1181, ORF 1183, ORF 1184, ORF 1186, ORF 1187, ORF 1188, ORF 1192, ORF 1194, ORF 1197, and their representative fragments.

5 Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a Chlamydia trachomatis polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, such as for example triose phosphate isomerase or pyruvate kinase, and in that they comprise a nucleotide sequence chosen from the following sequences: 10 ORF10; ORF44; ORF45; ORF46; ORF47; ORF93; ORF101; ORF102; ORF103; ORF106; ORF107; ORF120; ORF121; ORF130; ORF135; ORF140; ORF143; ORF144; ORF145; ORF158; ORF159; ORF160; ORF161; ORF192; ORF193; ORF196; ORF197; ORF198; ORF199; ORF227; ORF229; ORF236; ORF236; ORF239; ORF243; ORF245; ORF265; ORF297; ORF331; ORF333; ORF359; ORF360; ORF374; ORF404; ORF405; ORF405; ORF410; ORF415; ORF415; ORF416; 15 ORF417; ORF432; ORF460; ORF461; ORF462; ORF495; ORF513; ORF515; ORF566; ORF566; ORF566; ORF689; ORF613; ORF645; ORF646; ORF652; ORF653; ORF654; ORF672; ORF673; ORF674; ORF682; ORF684; ORF692; ORF700; ORF725; ORF801; ORF802; ORF835; ORF836; ORF860; ORF861; ORF862; ORF863; ORF869; ORF869; ORF925; ORF964; ORF983 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, such as for example CTP synthetase or GMP synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF142; ORF142; ORF169; ORF256; ORF268; ORF325; ORF352; ORF366; ORF435; ORF444; ORF528; ORF529; ORF530; ORF548; ORF549; ORF601; ORF602; ORF617; ORF619; ORF644; ORF745; ORF971; ORF972; ORF1023 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, such as for example DNA polymerases or DNA topoisomerases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF5; ORF12; ORF82; ORF96; ORF97; ORF98; ORF99; ORF100; ORF105; ORF118; ORF136; ORF137; ORF163; ORF190; ORF204; ORF259; ORF260; ORF262; ORF290; ORF300; ORF301; ORF302; ORF387; ORF427; ORF434; ORF441; ORF444; ORF471; ORF595; ORF596; ORF597;

ORF599; ORF600; ORF605; ORF612; ORF624; ORF625; ORF650; ORF657; ORF658; ORF702; ORF703; ORF704; ORF708; ORF719; ORF766; ORF767; ORF775; ORF779; ORF787; ORF788; ORF794; ORF841; ORF842; ORF883; ORF884; ORF907; ORF918; ORF924; ORF928; ORF929; ORF962; ORF962; ORF963; ORF969; ORF970; ORF975; ORF979; ORF995; ORF1031; ORF1032 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the

invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, such as for example serine hydroxymethyl transferase or the proteins which load amino acids onto transfer RNAs, and in that they comprise a nucleotide sequence chosen from the following sequences: ORF27; ORF41; ORF55; ORF56; ORF57; ORF59; ORF62; ORF63; ORF64; ORF65; ORF119; ORF132; ORF240; ORF241; ORF277; ORF278; ORF279; ORF382; ORF406; ORF428; ORF442; ORF446; ORF447; ORF453; ORF454; ORF541; ORF542; ORF591; ORF608; ORF609; ORF610; ORF618; ORF648; ORF649; ORF660; ORF661; ORF677; ORF717; ORF765; ORF797; ORF871; ORF875; ORF920; ORF922; ORF937; ORF998; ORF1020; ORF1021; ORF1034; ORF1044;

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, such as for example protein kinases or proteases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF21; ORF22; ORF23; ORF24; ORF25; ORF26; ORF75; ORF84; ORF86; ORF92; ORF133; ORF151; ORF152; ORF157; ORF179; ORF209; ORF307; ORF326; ORF343; ORF344; ORF345; ORF371; ORF429; ORF519; ORF557; ORF586; ORF587; ORF630; ORF656; ORF706; ORF707; ORF730; ORF751; ORF752; ORF786; ORF847; ORF885; ORF923; ORF978; ORF1039; ORF1048 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, such as for example succinyl-CoA-synthesizing proteins or phosphatidylserine synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF4; ORF15; ORF16; ORF141; ORF173; ORF205; ORF205; ORF206; ORF207; ORF208; ORF312; ORF355; ORF415; ORF550; ORF558; ORF560; ORF561; ORF574; ORF574; ORF577; ORF578; ORF590; ORF614; ORF772; ORF808; ORF809; ORF904; ORF905; ORF905; ORF933;

35 ORF934; ORF934; ORF936 and one of their representative fragments.

ORF1046; ORF1049 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the synthesis of the wall, such as for example KDO transferase, and the proteins responsible for the attachment of certain sugars onto the exposed proteins, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF87; ORF196; ORF242; ORF269; ORF628; ORF629; ORF634; ORF635; ORF637; ORF638; ORF1019 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, such as for example initiation factors, RNA polymerases or certain chaperone proteins, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF112; ORF113; ORF332; ORF212; ORF213; ORF350; ORF362; ORF363; ORF364; ORF407; ORF451; ORF546; ORF643; ORF744; ORF746; ORF833; ORF868; ORF981; ORF982; ORF1003; ORF1011; ORF1042 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* ribosomal polypeptide or one of its representative fragments, such as for example the ribosomal proteins L21, L27 and S10, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF114; ORF115; ORF116; ORF328; ORF361; ORF375; ORF445; ORF543; ORF584; ORF585; ORF743; ORF813; ORF941; ORF942; ORF944; ORF946; ORF947; ORF948; ORF950; ORF951; ORF952; ORF953; ORF954; ORF955; ORF955; ORF957; ORF958; ORF960; ORF961; ORF1040; ORF1041; ORF1043; ORF1063; ORF1064 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transport polypeptide or one of its representative fragments, such as for example the proteins for transporting amino acids, sugars and certain oligopeptides, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF6; ORF50; ORF51; ORF80; ORF125; ORF126; ORF128; ORF129; ORF215; ORF246; ORF248; ORF249; ORF251; ORF252; ORF253; ORF255; ORF271; ORF275; ORF293; ORF309; ORF323; ORF324; ORF398; ORF401; ORF449; ORF511; ORF512; ORF564; ORF565; ORF667; ORF679; ORF680; ORF711; ORF712; ORF713; ORF714; ORF715; ORF730; ORF731; ORF736; ORF737; ORF738; ORF870; ORF908; ORF919; ORF977; ORF987; ORF988; ORF992; ORF993; ORF994; ORF1028; ORF1029 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the

invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the virulence process, such as for example the proteins analogous to the *Escherichia coli* vacB protein, and in that they comprise a nucleotide sequence chosen from the following sequences:

5 ORF20; ORF815; ORF816; ORF898; ORF1059; ORF1060 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the secretory system and/or which is secreted, such as for example proteins homologous to proteins in the secretory system of certain bacteria such as the Salmonellae or the Yersiniae, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF758; ORF889; ORF890; ORF891; ORF896; ORF897; ORF898 and one of their representative fragments.

Preferably, the invention also relates to nucleotide sequences according to the invention, characterized in that they encode a polypeptide specific to Chlamydiae or one of its representative fragments, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF22; ORF31; ORF32; ORF34; ORF35; ORF39; ORF40; ORF43; ORF48; ORF49; ORF50; ORF52; ORF53; ORF54; ORF72; ORF77; ORF78; ORF87; ORF90; ORF95; ORF108; 20 ORF110; ORF121; ORF122; ORF123; ORF124; ORF127; ORF138; ORF144; ORF146; ORF153; ORF155; ORF164; ORF166; ORF175; ORF182; ORF184; ORF186; ORF187; ORF188; ORF202; ORF210; ORF247; ORF258; ORF266; ORF267; ORF270; ORF273; ORF274; ORF295; ORF296; ORF305; ORF306; ORF309; ORF318; ORF319; ORF322; ORF326; ORF342; ORF357; ORF376; ORF379; ORF380; ORF388; ORF390; ORF400; ORF431; ORF433; ORF438; ORF443; ORF456; 25 ORF457; ORF458; ORF464; ORF468; ORF470; ORF473; ORF486; ORF489; ORF497; ORF501; ORF503; ORF504; ORF508; ORF512; ORF521; ORF522; ORF523; ORF524; ORF533; ORF535; ORF536; ORF537; ORF538; ORF539; ORF540; ORF554; ORF563; ORF572; ORF579; ORF595; ORF603; ORF604; ORF606; ORF607; ORF615; ORF616; ORF622; ORF641; ORF642; ORF659; ORF668; ORF670; ORF693; ORF695; ORF699; ORF703; ORF704; ORF716; ORF726; 30 ORF728; ORF739; ORF742; ORF747; ORF750; ORF751; ORF755; ORF757; ORF759; ORF761; ORF762; ORF763; ORF764; ORF773; ORF780; ORF781; ORF789; ORF800; ORF803; ORF804; ORF818; ORF820; ORF822; ORF823; ORF824; ORF827; ORF828; ORF839; ORF849; ORF850; ORF851; ORF852; ORF855; ORF856; ORF857; ORF858; ORF859; ORF860; ORF861; ORF862; ORF863; ORF865; ORF868; ORF869; ORF870; ORF871; ORF872; ORF873; ORF874; ORF875; 35 ORF877; ORF878; ORF880; ORF882; ORF884; ORF886; ORF893; ORF901; ORF906; ORF910; ORF912; ORF915; ORF916; ORF917; ORF926; ORF929; ORF933; ORF965; ORF967; ORF968; ORF984; ORF986; ORF989; ORF990; ORF996; ORF997; ORF1001; ORF1002; ORF1013; ORF1016; ORF1031; ORF1033; ORF1035; ORF1049; ORF1051; ORF1052; ORF1054; ORF1056; ORF1057; ORF1058; ORF1062; ORF1070; ORF1071; ORF1073 and one of their representative fragments.

Also forming part of the invention are polypeptides encoded by the polynucleotides of the invention, as well as fusion polypeptides comprising such polypeptides. In one embodiment, the polypeptides and fusion polypeptides immunoreact with seropositive serum of an individual infected with *Chlamydia trachomatis*. For example, described below, are polypeptide sequences exhibiting particularly preferable characteristics. For each group of preferred polypeptides described below, it is to be understood that in addition to the individual polypeptides listed, in instances wherein such polypeptides are encoded as part of «combined» ORFs, such «combined» polypeptides are also to be included within the preferred group.

The subject of the invention is also a polypeptide according to the invention, characterized in that it is a polypeptide of the cellular envelope, preferably of the outer cellular envelope, of *Chlamydia trachomatis* or one of its representative fragments. According to the invention, the said polypeptide is preferably chosen from the polypeptides having the following sequences:

SEQ ID No. 3; SEQ ID No. 19; SEQ ID No. 51; SEQ ID No. 189; SEQ ID No. 212; SEQ ID No. 213; 20 SEQ ID No. 324; SEQ ID No. 477; SEQ ID No. 478; SEQ ID No. 479; SEQ ID No. 481; SEQ ID No. 482; SEQ ID No. 483; SEQ ID No. 484; SEQ ID No. 486; SEQ ID No. 488; SEQ ID No. 489; SEQ ID No. 490; SEQ ID No. 572; SEQ ID No. 573; SEQ ID No. 742; SEQ ID No. 817; SEQ ID No. 818; SEQ ID No. 820; SEQ ID No. 1035; SEQ ID No. 1036; SEQ ID No. 1037; SEQ ID No. 1038; SEQ ID No. 1070; SEQ ID No. 1071; SEQ ID No. 1073 and 25 one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 2; SEQ ID No. 3; SEQ ID No. 5; SEQ ID No. 8; SEQ ID No. 9; SEQ ID No. 10;
SEQ ID No. 11; SEQ ID No. 12; SEQ ID No. 17; SEQ ID No. 21; SEQ ID No. 26; SEQ ID No. 27;
SEQ ID No. 28; SEQ ID No. 29; SEQ ID No. 30; SEQ ID No. 31; SEQ ID No. 33; SEQ ID No. 35;
SEQ ID No. 37; SEQ ID No. 39; SEQ ID No. 40; SEQ ID No. 41; SEQ ID No. 42; SEQ ID No. 43;
SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 46; SEQ ID No. 47; SEQ ID No. 48; SEQ ID No. 49;
SEQ ID No. 52; SEQ ID No. 53; SEQ ID No. 55; SEQ ID No. 56; SEQ ID No. 58; SEQ ID No. 65;

SEQ ID No. 66; SEQ ID No. 68; SEQ ID No. 70; SEQ ID No. 74; SEQ ID No. 75; SEQ ID No. 76; SEQ ID No. 78; SEQ ID No. 79; SEQ ID No. 81; SEQ ID No. 82; SEQ ID No. 83; SEQ ID No. 86; SEQ ID No. 91; SEQ ID No. 92; SEQ ID No. 94; SEQ ID No. 97; SEQ ID No. 100; SEQ ID No. 102; SEQ ID No. 103; SEQ ID No. 105; SEQ ID No. 106; SEQ ID No. 107; SEQ ID No. 109; 5 SEQ ID No. 110; SEQ ID No. 111; SEQ ID No. 112; SEQ ID No. 113; SEQ ID No. 114; SEQ ID No. 115; SEQ ID No. 116; SEQ ID No. 117; SEQ ID No. 120; SEQ ID No. 122; SEQ ID No. 123; SEQ ID No. 130; SEQ ID No. 134; SEQ ID No. 135; SEQ ID No. 137; SEQ ID No. 140; SEQ ID No. 141; SEQ ID No. 143; SEQ ID No. 144; SEQ ID No. 145; SEQ ID No. 147; SEQ ID No. 148; SEQ ID No. 149; SEQ ID No. 150; SEQ ID No. 151; 10 SEQ ID No. 155; SEQ ID No. 156; SEQ ID No. 162; SEQ ID No. 163; SEQ ID No. 164; SEQ ID No. 165; SEQ ID No. 166; SEQ ID No. 167; SEQ ID No. 168; SEQ ID No. 169; SEQ ID No. 170; SEQ ID No. 171; SEQ ID No. 173; SEQ ID No. 175; SEQ ID No. 176; SEQ ID No. 177; SEQ ID No. 183; SEQ ID No. 181; SEQ ID No. 184; SEQ ID No. 186; SEQ ID No. 187; SEQ ID No. 188; SEQ ID No. 190; SEQ ID No. 191; SEQ ID No. 192; 15 SEQ ID No. 194; SEQ ID No. 195; SEQ ID No. 196; SEQ ID No. 197; SEQ ID No. 198; SEQ ID No. 199; SEQ ID No. 201; SEQ ID No. 202; SEQ ID No. 204; SEQ ID No. 206; SEQ ID No. 207; SEQ ID No. 209; SEQ ID No. 212; SEQ ID No. 213; SEQ ID No. 217; SEQ ID No. 219; SEQ ID No. 220; SEQ ID No. 221; SEQ ID No. 222; SEQ ID No. 223; SEQ ID No. 224; SEQ ID No. 225; SEQ ID No. 227; SEQ ID No. 228; SEQ ID No. 231; 20 SEQ ID No. 232; SEQ ID No. 234; SEQ ID No. 236; SEQ ID No. 237; SEQ ID No. 243; SEQ ID No. 244; SEQ ID No. 245; SEQ ID No. 247; SEQ ID No. 248; SEQ ID No. 249; SEQ ID No. 252; SEQ ID No. 254; SEQ ID No. 257; SEQ ID No. 260; SEQ ID No. 261; SEQ ID No. 263; SEQ ID No. 265; SEQ ID No. 266; SEQ ID No. 267; SEQ ID No. 270; SEQ ID No. 271; SEQ ID No. 272; SEQ ID No. 274; SEQ ID No. 276; SEQ ID No. 277; 25 SEQ ID No. 278; SEQ ID No. 279; SEQ ID No. 282; SEQ ID No. 283; SEQ ID No. 284; SEQ ID No. 285; SEQ ID No. 287; SEQ ID No. 289; SEQ ID No. 290; SEQ ID No. 291; SEQ ID No. 294; SEQ ID No. 298; SEQ ID No. 305; SEQ ID No. 306; SEQ ID No. 310; SEQ ID No. 311; SEQ ID No. 313; SEQ ID No. 315; SEQ ID No. 316; SEQ ID No. 319; SEQ ID No. 320; SEQ ID No. 322; SEQ ID No. 323; SEQ ID No. 325; SEQ ID No. 326; 30 SEQ ID No. 327; SEQ ID No. 328; SEQ ID No. 330; SEQ ID No. 331; SEQ ID No. 332; SEQ ID No. 333; SEQ ID No. 334; SEQ ID No. 335; SEQ ID No. 336; SEQ ID No. 338; SEQ ID No. 339; SEQ ID No. 340; SEQ ID No. 341; SEQ ID No. 345; SEQ ID No. 344; SEQ ID No. 348; SEQ ID No. 349; SEQ ID No. 350; SEQ ID No. 351; SEQ ID No. 352; SEQ ID No. 353; SEQ ID No. 356; SEQ ID No. 357; SEQ ID No. 358; SEQ ID No. 361; 35 SEQ ID No. 362; SEQ ID No. 366; SEQ ID No. 367; SEQ ID No. 368; SEQ ID No. 370;

	SEQ ID No. 372;	SEQ ID No. 373;	SEQ ID No. 375;	SEQ ID No. 377;	SEQ ID No. 378;
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	SEQ ID No. 385;	SEQ ID No. 387;	SEQ ID No. 389;	SEQ ID No. 390;	SEQ ID No. 391;
	SEQ ID No. 393;	SEQ ID No. 396;	SEQ ID No. 398;	SEQ ID No. 399;	SEQ ID No. 403;
5	SEQ ID No. 404;	SEQ ID No. 406;	SEQ ID No. 407;	SEQ ID No. 413;	SEQ ID No. 414;
	SEQ ID No. 417;	SEQ ID No. 418;	SEQ ID No. 420;	SEQ ID No. 421;	SEQ ID No. 424;
	SEQ ID No. 426;	SEQ ID No. 427;	SEQ ID No. 428;	SEQ ID No. 430;	SEQ ID No. 433;
10	SEQ ID No. 434;	SEQ ID No. 435;	SEQ ID No. 436;	SEQ ID No. 437;	SEQ ID No. 440;
	SEQ ID No. 443;	SEQ ID No. 446;	SEQ ID No. 448;	SEQ ID No. 450;	SEQ ID No. 451;
	SEQ ID No. 454;	SEQ ID No. 455;	SEQ ID No. 457;	SEQ ID No. 458;	SEQ ID No. 459;
	SEQ ID No. 463;	SEQ ID No. 464;	SEQ ID No. 466;	SEQ ID No. 467;	SEQ ID No. 468;
	SEQ ID No. 469;	SEQ ID No. 470;	SEQ ID No. 473;	SEQ ID No. 474;	SEQ ID No. 475;
	SEQ ID No. 476;	SEQ ID No. 477;	SEQ ID No. 479;	SEQ ID No. 480;	SEQ ID No. 481;
	SEQ ID No. 483;	SEQ ID No. 484;	SEQ ID No. 485;	SEQ ID No. 486;	SEQ ID No. 487;
15	SEQ ID No. 488;	SEQ ID No. 491;	SEQ ID No. 493;	SEQ ID No. 496;	SEQ ID No. 497;
	SEQ ID No. 498;	SEQ ID No. 500;	SEQ ID No. 501;	SEQ ID No. 503;	SEQ ID No. 504;
	SEQ ID No. 508;	SEQ ID No. 512;	SEQ ID No. 513;	SEQ ID No. 514;	SEQ ID No. 519;
	SEQ ID No. 521;	SEQ ID No. 523;	SEQ ID No. 524;	SEQ ID No. 526;	SEQ ID No. 527;
	SEQ ID No. 529;	SEQ ID No. 530;	SEQ ID No. 531;	SEQ ID No. 532;	SEQ ID No. 534;
20	SEQ ID No. 536;	SEQ ID No. 537;	SEQ ID No. 538;	SEQ ID No. 540;	SEQ ID No. 541;
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	SEQ ID No. 547;	SEQ ID No. 551;	SEQ ID No. 552;	SEQ ID No. 553;	SEQ ID No. 555;
	SEQ ID No. 558;	SEQ ID No. 559;	SEQ ID No. 560;	SEQ ID No. 561;	SEQ ID No. 562;
	SEQ ID No. 566;	SEQ ID No. 567;	SEQ ID No. 568;	SEQ ID No. 569;	SEQ ID No. 571;
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	SEQ ID No. 582;	SEQ ID No. 585;	SEQ ID No. 587;	SEQ ID No. 589;	SEQ ID No. 592;
	SEQ ID No. 593;	SEQ ID No. 595;	SEQ ID No. 596;	SEQ ID No. 597;	SEQ ID No. 599;
	SEQ ID No. 601;	SEQ ID No. 602;	SEQ ID No. 603;	SEQ ID No. 604;	SEQ ID No. 608;
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	SEQ ID No. 617;	SEQ ID No. 618;	SEQ ID No. 621;	SEQ ID No. 622;	SEQ ID No. 623;
	SEQ ID No. 624;	SEQ ID No. 625;	SEQ ID No. 628;	SEQ ID No. 632;	SEQ ID No. 633;
	SEQ ID No. 634;	SEQ ID No. 635;	SEQ ID No. 637;	SEQ ID No. 638;	SEQ ID No. 640;
	SEQ ID No. 641;	SEQ ID No. 643;	SEQ ID No. 646;	SEQ ID No. 648;	SEQ ID No. 649;
	SEQ ID No. 651;	SEQ ID No. 652;	SEQ ID No. 653;	SEQ ID No. 654;	SEQ ID No. 655;
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	SEQ ID No. 669;	SEQ ID No. 670;	SEQ ID No. 671;	SEQ ID No. 672;	SEQ ID No. 673;
	SEQ ID No. 674;	SEQ ID No. 676;	SEQ ID No. 677;	SEQ ID No. 678;	SEQ ID No. 680;
	SEQ ID No. 682;	SEQ ID No. 683;	SEQ ID No. 684;	SEQ ID No. 686;	SEQ ID No. 688;
	SEQ ID No. 689;	SEQ ID No. 690;	SEQ ID No. 691;	SEQ ID No. 692;	SEQ ID No. 693;
5	SEQ ID No. 695;	SEQ ID No. 696;	SEQ ID No. 698;	SEQ ID No. 701;	SEQ ID No. 703;
	SEQ ID No. 704;	SEQ ID No. 705;	SEQ ID No. 706;	SEQ ID No. 707;	SEQ ID No. 709;
	SEQ ID No. 710;	SEQ ID No. 711;	SEQ ID No. 712;	SEQ ID No. 713;	SEQ ID No. 714;
	SEQ ID No. 715;	SEQ ID No. 717;	SEQ ID No. 718;	SEQ ID No. 720;	SEQ ID No. 721;
	SEQ ID No. 722;	SEQ ID No. 724;	SEQ ID No. 726;	SEQ ID No. 728;	SEQ ID No. 729;
10	SEQ ID No. 730;	SEQ ID No. 731;	SEQ ID No. 732;	SEQ ID No. 733;	SEQ ID No. 734;
	SEQ ID No. 737;	SEQ ID No. 738;	SEQ ID No. 739;	SEQ ID No. 740;	SEQ ID No. 742;
	SEQ ID No. 743;	SEQ ID No. 744;	SEQ ID No. 745;	SEQ ID No. 746;	SEQ ID No. 748;
	SEQ ID No. 750;	SEQ ID No. 751;	SEQ ID No. 752;	SEQ ID No. 753;	SEQ ID No. 754;
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15	SEQ ID No. 764;	SEQ ID No. 766;	SEQ ID No. 768;	SEQ ID No. 769;	SEQ ID No. 771;
	SEQ ID No. 772;	SEQ ID No. 773;	SEQ ID No. 774;	SEQ ID No. 775;	SEQ ID No. 776;
	SEQ ID No. 777;	SEQ ID No. 778;	SEQ ID No. 779;	SEQ ID No. 780;	SEQ ID No. 781;
	SEQ ID No. 782;	SEQ ID No. 783;	SEQ ID No. 786;	SEQ ID No. 787;	SEQ ID No. 788;
	SEQ ID No. 789;	SEQ ID No. 790;	SEQ ID No. 793;	SEQ ID No. 798;	SEQ ID No. 800;
20	SEQ ID No. 802;	SEQ ID No. 803;	SEQ ID No. 806;	SEQ ID No. 808;	SEQ ID No. 809;
	SEQ ID No. 810;	SEQ ID No. 811;	SEQ ID No. 813;	SEQ ID No. 814;	SEQ ID No. 817;
	SEQ ID No. 820;	SEQ ID No. 822;	SEQ ID No. 824;	SEQ ID No. 825;	SEQ ID No. 827;
	SEQ ID No. 828;	SEQ ID No. 829;	SEQ ID No. 830;	SEQ ID No. 833;	SEQ ID No. 834;
	SEQ ID No. 835;	SEQ ID No. 837;	SEQ ID No. 838;	SEQ ID No. 839;	SEQ ID No. 840;
25	SEQ ID No. 841;	SEQ ID No. 842;	SEQ ID No. 843;	SEQ ID No. 845;	SEQ ID No. 848;
	SEQ ID No. 849;	SEQ ID No. 850;	SEQ ID No. 851;	SEQ ID No. 852;	SEQ ID No. 854;
	SEQ ID No. 855;	SEQ ID No. 856;	SEQ ID No. 857;	SEQ ID No. 859;	SEQ ID No. 860;
	SEQ ID No. 862;	SEQ ID No. 863;	SEQ ID No. 864;	SEQ ID No. 866;	SEQ ID No. 869;
	SEQ ID No. 872;	SEQ ID No. 873;	SEQ ID No. 874;	SEQ ID No. 878;	SEQ ID No. 879;
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	SEQ ID No. 886;	SEQ ID No. 887;	SEQ ID No. 892;	SEQ ID No. 893;	SEQ ID No. 894;
	SEQ ID No. 895;	SEQ ID No. 897;	SEQ ID No. 899;	SEQ ID No. 900;	SEQ ID No. 901;
	SEQ ID No. 904;	SEQ ID No. 906;	SEQ ID No. 909;	SEQ ID No. 910;	SEQ ID No. 912;
	SEQ ID No. 914;	SEQ ID No. 917;	SEQ ID No. 920;	SEQ ID No. 921;	SEQ ID No. 922;
35	SEQ ID No. 923;	SEQ ID No. 924;	SEQ ID No. 925;	SEQ ID No. 926;	SEQ ID No. 927;

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                                          SEQ ID No. 934;
                                                            SEQ ID No. 935;
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                       SEQ ID No. 940;
                                         SEQ ID No. 941;
                                                            SEQ ID No. 942;
                                                                              SEQ ID No. 943;
    SEQ ID No. 944;
                       SEQ ID No. 945;
                                         SEQ ID No. 947;
                                                            SEQ ID No. 948;
                                                                              SEQ ID No. 951;
    SEQ ID No. 952;
                       SEQ ID No. 953;
                                         SEQ ID No. 954;
                                                            SEQ ID No. 955;
                                                                              SEQ ID No. 956;
    SEQ ID No. 957;
                       SEQ ID No. 958;
                                         SEQ ID No. 960;
                                                            SEQ ID No. 961;
                                                                              SEQ ID No. 962;
    SEQ ID No. 963;
                       SEQ ID No. 964;
                                         SEQ ID No. 966;
                                                            SEQ ID No. 967;
                                                                              SEQ ID No. 969;
    SEQ ID No. 970;
                       SEQ ID No. 971;
                                         SEQ ID No. 973;
                                                            SEQ ID No. 974;
                                                                              SEQ ID No. 979;
    SEQ ID No. 980;
                       SEQ ID No. 981;
                                                            SEQ ID No. 984;
                                         SEQ ID No. 982;
                                                                              SEQ ID No. 988;
    SEQ ID No. 989;
                       SEQ ID No. 990;
                                         SEQ ID No. 991;
                                                            SEQ ID No. 995;
                                                                              SEQ ID No. 996;
    SEQ ID No. 999;
                      SEQ ID No. 1001;
                                        SEQ ID No. 1003;
                                                           SEQ ID No. 1004;
                                                                             SEQ ID No. 1005;
    SEQ ID No. 1006;
                       SEQ ID No. 1007;
                                         SEQ ID No. 1009;
                                                           SEQ ID No. 1010;
                                                                             SEQ ID No. 1011;
    SEQ ID No. 1012;
                       SEQ ID No. 1013;
                                         SEQ ID No. 1014;
                                                           SEQ ID No. 1016;
                                                                             SEQ ID No. 1017;
    SEQ ID No. 1018;
                      SEQ ID No. 1020;
                                         SEQ ID No. 1021;
                                                           SEQ ID No. 1025;
                                                                             SEQ ID No. 1026;
    SEQ ID No. 1027;
                      SEQ ID No. 1029;
                                         SEQ ID No. 1030;
                                                           SEQ ID No. 1031;
                                                                             SEQ ID No. 1035;
    SEQ ID No. 1036;
                      SEQ ID No. 1037;
                                        SEQ ID No. 1038;
                                                           SEQ ID No. 1039;
                                                                             SEQ ID No. 1040;
    SEQ ID No. 1044;
                      SEQ ID No. 1045;
                                        SEQ ID No. 1047;
                                                          SEQ ID No. 1048;
                                                                            SEQ ID No. 1050;
    SEQ ID No. 1051; SEQ ID No. 1052;
                                        SEQ ID No. 1053;
                                                          SEQ ID No. 1055;
                                                                            SEQ ID No. 1056;
    SEQ ID No. 1057;
                      SEQ ID No. 1058;
                                        SEQ ID No. 1061;
                                                          SEQ ID No. 1062; SEQ ID No. 1063;
    SEQ ID No. 1064; SEQ ID No. 1065; SEQ ID No. 1066; SEQ ID No. 1068; SEQ ID No. 1069;
20
    SEQ ID No. 1072; SEQ ID No. 1074; SEQ ID No. 1076 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 4 and 6 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

25 SEQ ID No. 7; SEQ ID No. 14; SEQ ID No. 16; SEQ ID No. 32; SEQ ID No. 34; SEQ ID No. 36; SEQ ID No. 38; SEQ ID No. 50; SEQ ID No. 57; SEQ ID No. 59; SEQ ID No. 61; SEQ ID No. 62; SEQ ID No. 63; SEQ ID No. 64; SEQ ID No. 67; SEQ ID No. 69; SEQ ID No. 72; SEQ ID No. 77; SEQ ID No. 80; SEQ ID No. 84; SEQ ID No. 87; SEQ ID No. 93; SEQ ID No. 95; SEQ ID No. 99; SEQ ID No. 108; SEQ ID No. 119; SEQ ID No. 125; SEQ ID No. 126; SEQ ID No. 129; 30 SEQ ID No. 131; SEQ ID No. 136; SEQ ID No. 139; SEQ ID No. 146; SEQ ID No. 152; SEQ ID No. 154; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 172; SEQ ID No. 179; SEQ ID No. 182; SEQ ID No. 185; SEQ ID No. 200; SEQ ID No. 203; SEQ ID No. 205; SEQ ID No. 239; SEQ ID No. 242; SEQ ID No. 250; SEQ ID No. 253; SEQ ID No. 256; SEQ ID No. 259; SEQ ID No. 262; SEQ ID No. 268; SEQ ID No. 275; SEQ ID No. 281; SEQ ID No. 286; 35 SEQ ID No. 288; SEQ ID No. 292; SEQ ID No. 295; SEQ ID No. 296;

	SEQ ID No. 297;	SEQ ID No. 299;	SEQ ID No. 300;	SEQ ID No. 308;	SEQ ID No. 314;
	SEQ ID No. 317;	SEQ ID No. 318;	SEQ ID No. 324;	SEQ ID No. 342;	SEQ ID No. 343;
	SEQ ID No. 355;	SEQ ID No. 360;	SEQ ID No. 374;	SEQ ID No. 376;	SEQ ID No. 386;
	SEQ ID No. 388;	SEQ ID No. 392;	SEQ ID No. 394;	SEQ ID No. 395;	SEQ ID No. 402;
5	SEQ ID No. 405;	SEQ ID No. 411;	SEQ ID No. 415;	SEQ ID No. 416;	SEQ ID No. 422;
	SEQ ID No. 423;	SEQ ID No. 429;	SEQ ID No. 432;	SEQ ID No. 441;	SEQ ID No. 442;
	SEQ ID No. 444;	SEQ ID No. 449;	SEQ ID No. 452;	SEQ ID No. 456;	SEQ ID No. 460;
	SEQ ID No. 461;	SEQ ID No. 465;	SEQ ID No. 471;	SEQ ID No. 472;	SEQ ID No. 482;
	SEQ ID No. 489;	SEQ ID No. 492;	SEQ ID No. 494;	SEQ ID No. 495;	SEQ ID No. 502;
10	SEQ ID No. 505;	SEQ ID No. 506;	SEQ ID No. 509;	SEQ ID No. 516;	SEQ ID No. 517;
	SEQ ID No. 520;	SEQ ID No. 525;	SEQ ID No. 533;	SEQ ID No. 539;	SEQ ID No. 549;
	SEQ ID No. 554;	SEQ ID No. 557;	SEQ ID No. 563;	SEQ ID No. 570;	SEQ ID No. 573;
	SEQ ID No. 581;	SEQ ID No. 590;	SEQ ID No. 591;	SEQ ID No. 600;	SEQ ID No. 607;
	SEQ ID No. 612;	SEQ ID No. 613;	SEQ ID No. 620;	SEQ ID No. 626;	SEQ ID No. 629;
15	SEQ ID No. 630;	SEQ ID No. 639;	SEQ ID No. 644;	SEQ ID No. 647;	SEQ ID No. 656;
	SEQ ID No. 659;	SEQ ID No. 661;	SEQ ID No. 685;	SEQ ID No. 687;	SEQ ID No. 699;
	SEQ ID No. 700;	SEQ ID No. 708;	SEQ ID No. 716;	SEQ ID No. 719;	SEQ ID No. 725;
	SEQ ID No. 747;	SEQ ID No. 749;	SEQ ID No. 756;	SEQ ID No. 765;	SEQ ID No. 767;
	SEQ ID No. 794;	SEQ ID No. 796;	SEQ ID No. 797;	SEQ ID No. 799;	SEQ ID No. 801;
20	SEQ ID No. 807;	SEQ ID No. 821;	SEQ ID No. 823;	SEQ ID No. 826;	SEQ ID No. 847;
	SEQ ID No. 853;	SEQ ID No. 861;	SEQ ID No. 870;	SEQ ID No. 871;	SEQ ID No. 875;
	SEQ ID No. 882;	SEQ ID No. 888;	SEQ ID No. 889;	SEQ ID No. 898;	SEQ ID No. 902;
	SEQ ID No. 903;	SEQ ID No. 911;	SEQ ID No. 916;	SEQ ID No. 931;	SEQ ID No. 939;
	SEQ ID No. 975;	SEQ ID No. 976;	SEQ ID No. 978;	SEQ ID No. 983;	SEQ ID No. 986;
25	SEQ ID No. 987;	SEQ ID No. 992;	SEQ ID No. 993;	SEQ ID No. 1000;	SEQ ID No. 1002;
	SEQ ID No. 1008;	SEQ ID No. 1019;	SEQ ID No. 1022;	SEQ ID No. 1032;	SEQ ID No. 1034;
	SEQ ID No. 1046;	SEQ ID No. 1054;	SEQ ID No. 1060;	SEQ ID No. 1071 a	and one of their
	representative fragr	nents.			

Preferably, the invention relates to a polypeptide according to the invention,

characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 4; SEQ ID No. 6; SEQ ID No. 13; SEQ ID No. 20; SEQ ID No. 51; SEQ ID No. 71; SEQ ID No. 88; SEQ ID No. 118; SEQ ID No. 128; SEQ ID No. 132; SEQ ID No. 133; SEQ ID No. 158; SEQ ID No. 159; SEQ ID No. 174;

SEQ ID No. 180; SEQ ID No. 189; SEQ ID No. 210; SEQ ID No. 211; SEQ ID No. 214;

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SEQ ID No. 215;
                        SEQ ID No. 226;
                                          SEQ ID No. 229;
                                                             SEQ ID No. 233;
                                                                               SEQ ID No. 235;
     SEQ ID No. 240;
                        SEQ ID No. 246;
                                          SEQ ID No. 251;
                                                             SEQ ID No. 255;
                                                                               SEQ ID No. 273;
     SEQ ID No. 354;
                        SEQ ID No. 364;
                                          SEQ ID No. 369;
                                                            SEQ ID No. 371;
                                                                               SEQ ID No. 397;
     SEQ ID No. 401;
                        SEQ ID No. 409;
                                          SEO ID No. 412;
                                                            SEQ ID No. 419;
                                                                               SEQ ID No. 439;
 5 SEQ ID No. 453;
                       SEQ ID No. 462;
                                          SEQ ID No. 490;
                                                            SEQ ID No. 510;
                                                                               SEQ ID No. 511;
     SEQ ID No. 518;
                       SEQ ID No. 535;
                                          SEQ ID No. 548;
                                                            SEQ ID No. 550;
                                                                               SEQ ID No. 564;
     SEQ ID No. 565;
                       SEQ ID No. 578;
                                          SEQ ID No. 579;
                                                            SEQ ID No. 614;
                                                                               SEQ ID No. 631;
     SEQ ID No. 636;
                       SEQ ID No. 650;
                                          SEQ ID No. 662;
                                                            SEQ ID No. 667;
                                                                               SEQ ID No. 679;
     SEQ ID No. 681;
                       SEQ ID No. 702;
                                          SEQ ID No. 727;
                                                            SEQ ID No. 741;
                                                                               SEQ ID No. 763;
10 SEQ ID No. 791;
                       SEQ ID No. 792;
                                          SEQ ID No. 815;
                                                            SEQ ID No. 816;
                                                                               SEQ ID No. 832;
     SEQ ID No. 846;
                       SEQ ID No. 858;
                                          SEQ ID No. 865;
                                                            SEQ ID No. 867;
                                                                               SEQ ID No. 868;
    SEQ ID No. 877;
                       SEQ ID No. 891;
                                          SEQ ID No. 896;
                                                            SEQ ID No. 907;
                                                                               SEQ ID No. 908;
    SEQ ID No. 918;
                       SEQ ID No. 919;
                                          SEQ ID No. 932;
                                                            SEQ ID No. 959;
                                                                               SEQ ID No. 977;
     SEQ ID No. 994;
                      SEQ ID No. 998;
                                        SEQ ID No. 1024;
                                                          SEQ ID No. 1028;
                                                                             SEQ ID No. 1042;
15 SEQ ID No. 1067; SEQ ID No. 1070; SEQ ID No. 1073 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* surface exposed polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 2, SEQ ID No. 3, SEQ ID No. 21, SEQ ID No. 22, SEQ ID No. 23, SEQ ID No. 53, SEQ ID No. 77, SEQ ID No. 187, SEQ ID No. 203, SEQ ID No. 383, SEQ ID No. 477, SEQ ID No. 478, SEQ ID No. 479, SEQ ID No. 481, SEQ ID No. 482, SEQ ID No. 483, SEQ ID No. 484, SEQ ID No. 485, SEQ ID No. 486, SEQ ID No. 487, SEQ ID No. 488, SEQ ID No. 489, SEQ ID No. 490, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 593, SEQ ID No. 670, SEQ ID No. 693, SEQ ID No. 742, SEQ ID No. 749, SEQ ID No. 801, SEQ ID No. 817, SEQ ID No. 818, SEQ ID No. 819, SEQ ID No. 820, SEQ ID No. 851, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 1035, SEQ ID No. 1036, SEQ ID No. 1037, SEQ ID No. 1038, SEQ ID No. 1069, SEQ ID No. 1070, SEQ ID No. 1071, SEQ ID No. 1073, SEQ ID No. 1076, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1141, SEQ ID No. 1181, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* lipoprotein or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 29, SEQ ID No. 42, SEQ ID No. 66, SEQ ID No. 72, SEQ ID No. 76, SEQ ID No. 78, SEQ ID No. 148, SEQ ID No. 154, SEQ ID No. 180, SEQ ID No. 182, SEQ ID No. 184, SEQ ID No. 187, SEQ ID No. 200, SEQ ID No. 242, SEQ ID No. 245, SEQ ID No. 250, SEQ ID No. 253, SEQ ID No. 272, SEQ ID No. 274, SEQ ID No. 275, SEQ ID No. 308, SEQ ID No. 350, SEQ ID No. 362,

SEQ ID No. 383, SEQ ID No. 394, SEQ ID No. 396, SEQ ID No. 399, SEQ ID No. 422, SEQ ID No. 488, SEQ ID No. 535, SEQ ID No. 568, SEQ ID No. 573, SEQ ID No. 578, SEQ ID No. 593, SEQ ID No. 607, SEQ ID No. 625, SEQ ID No. 662, SEQ ID No. 669, SEQ ID No. 688, SEQ ID No. 690, SEQ ID No. 716, SEQ ID No. 773, SEQ ID No. 778, SEQ ID No. 781, SEQ ID No. 783, SEQ ID No. 788, SEQ ID No. 817, SEQ ID No. 848, SEQ ID No. 851, SEQ ID No. 853, SEQ ID No. 857, SEQ ID No. 875, SEQ ID No. 877, SEQ ID No. 886, SEQ ID No. 898, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 938, SEQ ID No. 976, SEQ ID No. 978, SEQ ID No. 990, SEQ ID No. 1005, SEQ ID No. 1021, SEQ ID No. 1035, SEQ ID No. 1069, SEQ ID No. 1083, SEQ ID No. 1088, SEQ ID No. 1089, SEQ ID No. 1091, SEQ ID No. 1092, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1100, SEQ ID No. 1105, SEQ ID No. 1108, SEQ ID No. 1117, SEQ ID No. 1120, SEQ ID No. 1121, SEQ ID No. 1140, SEQ ID No. 1157, SEQ ID No. 1159, SEQ ID No. 1163, SEQ ID No. 1165, SEQ ID No. 1167, SEQ ID No. 1169, SEQ ID No. 1161, SEQ ID No. 1173, SEQ ID No. 1174, SEQ ID No. 1177, SEQ ID No. 1180, SEQ ID No. 1181, SEQ ID No. 1186, SEQ ID No. 1194, SEQ ID No. 1197, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide involved in lipopolysaccharide (LPS) biosynthesis, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 17, SEQ ID No. 201, SEQ ID No. 691, SEQ ID No. 807, SEQ ID No. 936, SEQ ID No. 983, SEQ ID No. 1019, SEQ ID No. 1077, and their representative fragments.

Preferably, the invention relates to additional LPS-related polypeptides according to the invention, in that it is:

- (a) a Chlamydia trachomatis KDO (3-deoxy-D-manno-octylosonic acid)-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 41, SEQ ID No. 242, SEQ ID No. 269, SEQ ID No. 772, and one of their representative fragments;
 - (b) a *Chlamydia trachomatis* phosphomannomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequence: SEQ ID No. 139, and its representative fragments;
 - (c) a *Chlamydia trachomatis* phosphoglucomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequence: SEQ ID No. 567 and its representative fragments; and
- (d) a Chlamydia trachomatis lipid A component-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following
 sequences: SEQ ID No. 4, SEQ ID No. 933, SEQ ID No. 934, SEQ ID No. 935, SEQ ID No. 1185,

and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments that contains an RGD sequence and is also an outer membrane protein, and in that it is chosen from the polypeptides having the following sequences: SEQ. ID No. 488, SEQ ID No. 489, SEQ ID No. 571, SEQ ID No. 572, SEQ No. 573, SEQ ID No. 716 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments that is cysteine-rich and contains RGD sequence, and in that it is chosen from the polypeptides having the following sequence:

SEQ ID No. 144 and one of its representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* outer membrane polypeptide that contains cysteines in their first 30 amino acids and also contain an RGD sequence, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 101, SEQ ID No. 122, SEQ ID No. 308, SEQ ID No. 488, SEQ ID No. 489, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 651, SEQ ID No. 679, SEQ ID No. 680, SEQ ID No. 705, SEQ ID No. 716, SEQ ID No. 763, SEQ ID No. 870, SEQ ID No. 878, SEQ ID No. 879, SEQ ID No. 995, SEQ ID No. 1028, SEQ ID No. 1029, SEQ ID No. 1176, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments that contains RGD sequences homologous to *Chlamydia pneumoniae* polypeptides containing RGD sequences, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 28, SEQ ID No. 101, SEQ ID No. 125, SEQ ID No. 155, SEQ ID No. 156, SEQ ID No. 286, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 763, SEQ ID No. 870, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* Type III or non-Type III secreted polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 180, SEQ ID No. 181, SEQ ID No. 207, SEQ ID No. 208, SEQ ID No. 372, SEQ ID No. 391, SEQ ID No. 399, SEQ ID No. 477, SEQ ID No. 486, SEQ ID No. 749, SEQ ID No. 758, SEQ ID No. 819, SEQ ID No. 878, SEQ ID No. 888, SEQ ID No. 896, SEQ ID No. 897, SEQ ID No. 900, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 1015, SEQ ID No. 1018, SEQ ID No. 1059, SEQ ID No. 1060, SEQ ID No. 1069, SEQ ID No. 1071, SEQ ID No. 1073, SEQ ID No. 1076, SEQ ID No. 1189, and their representative fragments.

is a *Chlamydia trachomatis* cell wall anchored surface polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 662, SEQ ID No. 681, SEQ ID No. 1182, SEQ ID No. 1192, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments not found in Chlamydia pneumoniae (Blastp P>e-10) and in that it is chosen from the polypeptides having the following sequences: SEQ ID No.2, SEQ ID No. 18, SEQ ID No. 60, SEQ ID No. 66, SEQ ID No. 67, SEQ ID No.68, SEQ ID No. 69, SEQ ID No. 70, SEQ ID No. 81, SEQ ID No. 89, SEQ ID No. 107, SEQ ID 10 No.108, SEQ ID No. 109, SEQ ID No.134, SEQ ID No. 147, SEQ ID No.191, SEQ ID No. 194, SEQ ID No. 216, SEQ ID No. 217, SEQ ID No. 218, SEQ ID No. 219, SEQ ID No. 220, SEQ ID No. 221, SEQ ID No. 222, SEQ ID No. 222, SEQ ID No. 223, SEQ ID No. 224, SEQ ID No. 225, SEQ ID No. 228, SEQ ID No. 235, SEQ ID No.257, SEQ ID No. 276, SEQ ID No. 277, SEQ ID No. 278, SEQ ID No No. 279, SEQ ID No. 280, SEQ ID No. 281, SEQ ID No. 282, SEQ ID No. 283, SEQ ID No. 284, 15 SEQ ID No. 285, SEQ ID No. 289, SEQ ID No.291, SEQ ID No. 298, SEQ ID No. 284, SEQ ID No. 313, SEQ ID No. 314, SEQ ID No. 315, SEQ ID No. 316, SEQ ID No. 334, SEQ ID No. 335, SEQ ID No. 336, SEQ ID No. 337, SEQ ID No. 338, SEQ ID No. 339, SEQ ID No. 340, SEQ ID No. 381, SEQ ID No. 393, SEQ ID No. 413, SEQ ID No. 418, SEQ ID No. 419, SEQ ID No. 419, SEQ ID No. 420, SEQ ID No. 421, SEQ ID No. 422, SEQ ID No. 423, SEQ ID No. 436, SEQ ID No. 460, SEQ ID 20 No. 475, SEQ ID No. 476, SEQ ID No. 480, SEQ ID No. 485, SEQ ID No. 487, SEQ ID No.491, SEQ ID No. 492, SEQ ID No. 493, SEQ ID No. 494, SEQ ID No. 496, SEQ ID No. 500, SEQ ID No. 504, SEQ ID No. 514, SEQ ID No. 527, SEQ ID No. 559, SEQ ID No.569, SEQ ID No. 570, SEQ ID No. 575, SEQ ID No. 580, SEQ ID No. 582, SEQ ID No. 593, SEQ ID No. 598, SEQ ID No.632, SEQ ID No.640, SEQ ID No.651, SEQ ID No.671, SEQ ID No. 690, SEQ ID No. 694, ID No. 698, SEQ ID 25 No. 710, SEQ ID No. 722, SEQ ID No. 723, SEQ ID No. 724, SEQ ID No. 770, SEQ ID No. 771, SEQ ID No. 782, SEQ ID No. 783, SEQ ID No. 784, SEQ ID No. 790, SEQ ID No. 795, SEQ ID No. 798, SEQ ID No. 805, SEQ ID No. 810, SEQ ID No. 817, SEQ ID No. 829, SEQ ID No. 830, SEQ ID No. 864, SEQ ID No. 866, SEQ ID No. 876, SEQ ID No. 887, SEQ ID No. 892, SEQ ID No. 899, SEQ ID No. 913, SEQ ID No. 921, SEQ ID No. 933, SEQ ID No. 938, SEQ ID No. 949, SEQ ID No. 30 956, SEQ ID No. 1010, SEQ ID No. 1017, SEQ ID No. 1018, SEQ ID No. 1027, SEQ ID No. 1030, SEQ ID No. 1037, SEQ ID No. 1038, SEQ ID No. 1047, SEQ ID No. 1072, SEQ ID No. 1074, SEQ ID No. 1075, SEQ ID No. 1078, SEQ ID No. 1079, SEQ ID No. 1081, SEQ ID No. 1083, SEQ ID No. 1084, SEQ ID No. 1087, SEQ ID No. 1088, SEQ ID No. 1089, SEQ ID No. 1091, SEQ ID No. 1092, SEQ ID No. 1094, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1098, SEQ ID No. 1104, SEQ 35 ID No. 1105, SEQ ID No. 1106, SEQ ID No. 1108, SEQ ID No. 1110, SEQ ID No. 1114, SEQ ID No.

1115, SEQ ID No. 1116, SEQ ID No. 1117, SEQ ID No. 1119, SEQ ID No. 1128, SEQ ID No. 1132, SEQ ID No. 1133, SEQ ID No. 1135, SEQ ID No. 1136, SEQ ID No. 1139, SEQ ID No. 1140, SEQ ID No. 1141, SEQ ID No. 1142, SEQ ID No. 1144, SEQ ID No. 1148, SEQ ID No. 1151, SEQ ID No. 1155, SEQ ID No. 1157, SEQ ID No. 1159, SEQ ID No. 1161, SEQ ID No. 1162, SEQ ID No. 1165, SEQ ID No. 1166, SEQ ID No. 1167, SEQ ID No. 1168, SEQ ID No. 1169, SEQ ID No. 1171, SEQ ID No. 1172, SEQ ID No. 1173, SEQ ID No. 1174, SEQ ID No. 1175, SEQ ID No. 1176, SEQ ID No. 1177, SEQ ID No. 1178, SEQ ID No. 1180, SEQ ID No. 1181, SEQ ID No. 1183, SEQ ID No. 1184, SEQ ID No. 1186, SEQ ID No. 1187, SEQ ID No. 1188, SEQ ID No. 1192, SEQ ID No. 1194, SEQ ID No. 1197, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 10; SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 46; SEQ ID No. 47; SEQ ID No. 93; 15 SEQ ID No. 101; SEQ ID No. 102; SEQ ID No. 103; SEQ ID No. 106: SEQ ID No. 107; SEQ ID No. 120; SEQ ID No. 121; SEQ ID No. 130; SEQ ID No. 135; SEQ ID No. 140; SEQ ID No. 143; SEQ ID No. 144; SEQ ID No. 145; SEQ ID No. 158; SEQ ID No. 159; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 192; SEQ ID No. 193; SEQ ID No. 196; SEQ ID No. 197; SEQ ID No. 198; SEQ ID No. 199; SEQ ID No. 227; SEQ ID No. 229; 20 SEQ ID No. 236; SEQ ID No. 236; SEQ ID No. 239; SEQ ID No. 243; SEQ ID No. 245; SEQ ID No. 264; SEQ ID No. 265; SEQ ID No. 297; SEQ ID No. 331; SEQ ID No. 333; SEQ ID No. 359; SEQ ID No. 360; SEQ ID No. 374; SEQ ID No. 404; SEQ ID No. 405; SEQ ID No. 405; SEQ ID No. 410; SEQ ID No. 415; SEQ ID No. 415; SEQ ID No. 416; SEQ ID No. 417; SEQ ID No. 432; SEQ ID No. 460; SEQ ID No. 461; SEQ ID No. 462; 25 SEQ ID No. 495; SEQ ID No. 513; SEQ ID No. 515; SEQ ID No. 566; SEQ ID No. 566; SEQ ID No. 566; SEQ ID No. 589; SEQ ID No. 613; SEQ ID No. 645; SEQ ID No. 646; SEQ ID No. 647; SEQ ID No. 652; SEQ ID No. 653; SEQ ID No. 654; SEQ ID No. 672; SEQ ID No. 673; SEQ ID No. 674; SEQ ID No. 682; SEQ ID No. 684; SEQ ID No. 692; SEQ ID No. 700; SEQ ID No. 725: SEQ ID No. 801; SEQ ID No. 802; SEQ ID No. 835; 30 SEQ ID No. 836; SEQ ID No. 837; SEQ ID No. 860; SEQ ID No. 861; SEQ ID No. 862; SEQ ID No. 863; SEQ ID No. 869; SEQ ID No. 869; SEQ ID No. 925; SEQ ID No. 964; SEQ ID No. 983 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, and in that it is

chosen from the polypeptides having the following sequences:

```
SEQ ID No. 142;
                      SEQ ID No. 142;
                                         SEQ ID No. 169;
                                                           SEQ ID No. 256;
                                                                              SEQ ID No. 268;
   SEQ ID No. 325;
                      SEQ ID No. 352;
                                         SEQ ID No. 366;
                                                           SEQ ID No. 435;
                                                                              SEQ ID No. 444;
   SEQ ID No. 528;
                      SEQ ID No. 529;
                                         SEQ ID No. 530;
                                                           SEQ ID No. 548;
                                                                              SEQ ID No. 549;
5 SEQ ID No. 601;
                      SEQ ID No. 602;
                                         SEQ ID No. 617;
                                                           SEQ ID No. 619;
                                                                              SEQ ID No. 644;
   SEQ ID No. 745; SEQ ID No. 971; SEQ ID No. 972; SEQ ID No. 1023 and one of their representative
   fragments.
```

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, and in that it is chosen from the polypeptides having the following sequences:

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SEQ ID No. 5; SEQ ID No. 12; SEQ ID No. 82; SEQ ID No. 96; SEQ ID No. 97; SEQ ID No. 98;
     SEQ ID No. 99;
                       SEQ ID No. 100;
                                          SEQ ID No. 105;
                                                            SEQ ID No. 118;
                                                                               SEQ ID No. 136;
     SEQ ID No. 137;
                        SEQ ID No. 163;
                                          SEQ ID No. 190;
                                                            SEQ ID No. 204;
                                                                               SEQ ID No. 259;
15 SEQ ID No. 260;
                        SEQ ID No. 262;
                                          SEQ ID No. 290;
                                                            SEQ ID No. 300;
                                                                               SEQ ID No. 301;
     SEQ ID No. 302;
                        SEQ ID No. 387;
                                          SEQ ID No. 427;
                                                            SEQ ID No. 434;
                                                                               SEQ ID No. 441;
     SEQ ID No. 444;
                        SEQ ID No. 471;
                                          SEQ ID No. 595;
                                                            SEQ ID No. 596;
                                                                               SEQ ID No. 597;
     SEQ ID No. 599;
                       SEQ ID No. 600;
                                          SEQ ID No. 605;
                                                            SEQ ID No. 612;
                                                                               SEQ ID No. 624;
    SEQ ID No. 625;
                       SEQ ID No. 650;
                                          SEQ ID No. 657;
                                                            SEQ ID No. 658;
                                                                               SEQ ID No. 702;
20 SEQ ID No. 703;
                       SEQ ID No. 704;
                                          SEQ ID No. 708;
                                                            SEQ ID No. 719;
                                                                               SEQ ID No. 766;
    SEQ ID No. 767;
                       SEQ ID No. 775;
                                          SEQ ID No. 779;
                                                            SEQ ID No. 787;
                                                                               SEQ ID No. 788;
    SEQ ID No. 794;
                       SEQ ID No. 841;
                                          SEQ ID No. 842;
                                                            SEQ ID No. 883;
                                                                               SEQ ID No. 884;
    SEQ ID No. 907;
                       SEQ ID No. 918;
                                          SEQ ID No. 924;
                                                            SEQ ID No. 928;
                                                                               SEQ ID No. 929;
    SEQ ID No. 962;
                       SEQ ID No. 962;
                                          SEQ ID No. 963;
                                                            SEQ ID No. 969;
                                                                               SEQ ID No. 970;
25 SEQ ID No. 975; SEQ ID No. 979; SEQ ID No. 995; SEQ ID No. 1031; SEQ ID No. 1032 and one of
    their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, and in that it is chosen from the polypeptides having the following sequences:

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SEQ ID No. 27; SEQ ID No. 41; SEQ ID No. 55; SEQ ID No. 56; SEQ ID No. 57; SEQ ID No. 59;
    SEQ ID No. 62; SEQ ID No. 63; SEQ ID No. 64; SEQ ID No. 65; SEQ ID No. 119; SEQ ID No. 132;
    SEQ ID No. 240;
                       SEQ ID No. 241;
                                         SEQ ID No. 277;
                                                           SEQ ID No. 278;
                                                                              SEQ ID No. 279;
    SEQ ID No. 382;
                       SEQ ID No. 406;
                                         SEQ ID No. 428;
                                                           SEQ ID No. 442;
                                                                              SEQ ID No. 446;
35 SEQ ID No. 447;
                       SEQ ID No. 453;
                                         SEQ ID No. 454;
                                                           SEQ ID No. 541;
                                                                              SEQ ID No. 542;
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SEQ ID No. 591; SEQ ID No. 608; SEQ ID No. 609; SEQ ID No. 610; SEQ ID No. 618; SEQ ID No. 648; SEQ ID No. 660; SEQ ID No. 649; SEQ ID No. 661; SEQ ID No. 677; SEQ ID No. 717; SEQ ID No. 765; SEQ ID No. 797; SEQ ID No. 871; SEQ 1D No. 875; SEQ ID No. 920; SEQ ID No. 922; SEQ ID No. 937: SEQ ID No. 998; SEQ ID No. 1020; 5 SEQ ID No. 1021; SEQ ID No. 1034; SEQ ID No. 1044; SEQ ID No. 1046; SEQ ID No. 1049 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 21; SEQ ID No. 22; SEQ ID No. 23; SEQ ID No. 24; SEQ ID No. 25; SEQ ID No. 26; SEQ ID No. 75; SEQ ID No. 84; SEQ ID No. 86; SEQ ID No. 92; SEQ ID No. 133; SEQ ID No. 151; SEQ ID No. 152; SEQ ID No. 157; SEQ ID No. 209; SEQ ID No. 179; SEQ ID No. 307; SEQ ID No. 326; SEQ ID No. 343; SEQ ID No. 344; SEQ ID No. 345; SEQ ID No. 371; 15 SEQ ID No. 429; SEQ ID No. 519; SEQ ID No. 557; SEQ ID No. 586; SEQ ID No. 587; SEQ ID No. 630; SEQ ID No. 656; SEQ ID No. 706; SEQ ID No. 707; SEQ ID No. 730; SEQ ID No. 751: SEQ ID No. 752; SEQ ID No. 786; SEQ ID No. 847; SEQ ID No. 885; SEQ ID No. 923; SEQ ID No. 978; SEQ ID No. 1039; SEQ ID No. 1048 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 4; SEQ ID No. 15; SEQ ID No. 16; SEQ ID No. 141; SEQ ID No. 173; SEQ ID No. 205; 25 SEQ ID No. 205; SEQ ID No. 206; **SEQ ID No. 207**; SEQ ID No. 208; SEQ ID No. 312; SEQ ID No. 355; SEQ ID No. 415; SEQ ID No. 550; SEQ ID No. 558; SEQ ID No. 560; SEQ ID No. 561; SEQ ID No. 574; SEQ ID No. 574; SEQ ID No. 577; SEQ ID No. 578; SEQ ID No. 590; SEQ ID No. 614; SEQ ID No. 772; SEQ ID No. 808; SEQ ID No. 809; SEQ ID No. 904; SEQ ID No. 905; SEQ ID No. 905; SEQ ID No. 933; SEQ ID No. 934; SEQ ID No. 934; SEQ ID No. 936 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the synthesis of the wall, and in that it is chosen from the polypeptides having the following sequences:

35 SEQ ID No. 87; SEQ ID No. 196; SEQ ID No. 242; SEQ ID No. 269; SEQ ID No. 628;

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SEQ ID No. 629; SEQ ID No. 634; SEQ ID No. 635; SEQ ID No. 637; SEQ ID No. 638; SEQ ID No. 1019 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, and in that it is chosen from the polypeptides having the following sequences:

```
SEQ ID No. 112;
                       SEQ ID No. 113;
                                          SEQ ID No. 332;
                                                             SEQ ID No. 212;
                                                                                SEQ ID No. 213;
    SEQ ID No. 350;
                       SEQ ID No. 362;
                                          SEQ ID No. 363;
                                                             SEQ ID No. 364;
                                                                                SEQ ID No. 407;
    SEQ ID No. 451;
                       SEQ ID No. 546;
                                          SEQ ID No. 643;
                                                            SEQ ID No. 744;
                                                                                SEQ ID No. 746;
10 SEQ ID No. 833;
                       SEQ ID No. 868;
                                         SEQ ID No. 981;
                                                            SEQ ID No. 982;
                                                                              SEQ ID No. 1003;
    SEQ ID No. 1011; SEQ ID No. 1042 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* ribosomal polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

```
15 SEQ ID No. 114;
                       SEQ ID No. 115;
                                         SEQ ID No. 116;
                                                           SEQ ID No. 328;
                                                                              SEQ ID No. 361;
    SEQ ID No. 375;
                       SEQ ID No. 445;
                                         SEQ ID No. 543;
                                                           SEQ ID No. 584;
                                                                              SEQ ID No. 585;
    SEQ ID No. 743;
                       SEQ ID No. 813;
                                         SEQ ID No. 941;
                                                           SEQ ID No. 942;
                                                                              SEQ ID No. 944;
    SEQ ID No. 946;
                       SEQ ID No. 947;
                                         SEQ ID No. 948;
                                                           SEQ ID No. 950;
                                                                              SEQ ID No. 951;
    SEQ ID No. 952;
                       SEQ ID No. 953;
                                         SEQ ID No. 954;
                                                           SEQ ID No. 955;
                                                                              SEQ ID No. 955;
20 SEQ ID No. 957;
                      SEQ ID No. 958;
                                         SEQ ID No. 960;
                                                          SEQ ID No. 961;
                                                                            SEQ ID No. 1040;
    SEQ ID No. 1041; SEQ ID No. 1043;
                                       SEQ ID No. 1063; SEQ ID No. 1064 and one of their
    fragments.
```

Preferably, the invention also relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transport polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

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SEQ ID No. 6; SEQ ID No. 50; SEQ ID No. 51; SEQ ID No. 80; SEQ ID No. 125; SEQ ID No. 126;
    SEQ ID No. 128;
                       SEQ ID No. 129;
                                          SEQ ID No. 215;
                                                            SEQ ID No. 246;
                                                                               SEQ ID No. 248;
    SEQ ID No. 249;
                       SEQ ID No. 251;
                                          SEQ ID No. 252;
                                                                               SEQ ID No. 255;
                                                            SEQ ID No. 253;
    SEQ ID No. 271;
                       SEQ ID No. 275;
                                          SEQ ID No. 293;
                                                                               SEQ ID No. 323;
                                                            SEQ ID No. 309;
    SEQ ID No. 324;
                       SEQ ID No. 398;
                                          SEQ ID No. 401;
                                                            SEQ ID No. 449;
                                                                               SEQ ID No. 511;
    SEQ ID No. 512;
                       SEQ ID No. 564;
                                          SEQ ID No. 565;
                                                            SEQ ID No. 667;
                                                                               SEQ ID No. 679;
    SEQ ID No. 680;
                       SEQ ID No. 711;
                                          SEQ ID No. 712;
                                                            SEQ ID No. 713;
                                                                               SEQ ID No. 714;
    SEQ ID No. 715;
                       SEQ ID No. 730;
                                          SEQ ID No. 731;
                                                            SEQ ID No. 736;
                                                                               SEQ ID No. 737;
    SEQ ID No. 738;
                       SEQ ID No. 870;
                                          SEQ ID No. 908;
                                                            SEQ ID No. 919;
                                                                               SEQ ID No. 977;
35
    SEQ ID No. 987;
                       SEQ ID No. 988;
                                          SEQ ID No. 992;
                                                            SEQ ID No. 993;
                                                                               SEQ ID No. 994;
```

35 SEQ ID No. 433;

SEQ ID No. 438;

SEQ ID No. 1028; SEQ ID No. 1029 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the virulence process, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 20; SEQ ID No. 815; SEQ ID No. 816; SEQ ID No. 898; SEQ ID No. 1059; SEQ ID No. 1060 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the secretory system and/or which is secreted, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 758; SEQ ID No. 888; SEQ ID No. 889; SEQ ID No. 890; SEQ ID No. 891; SEQ ID No. 896; SEQ ID No. 897; SEQ ID No. 898 and one of their representative fragments.

The secreted polypeptides, including the Type III and other, non-Type III secreted polypeptides, of the present invention, as well as the corresponding nucleotide sequences, may be detected by techniques known to persons skilled in the art, such as for example the techniques using cloning combined with vectors allowing the expression of the said polypeptides fused to export markers such as the *luc* gene for luciferase or the *PhoA* gene for alkaline phosphatase.

Preferably, the invention relates to a polypeptide according to the invention,

characterized in that it is a polypeptide specific to Chlamydiae or one of its representative fragments,
and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 22; SEQ ID No. 29; SEQ ID No. 31; SEQ ID No. 32; SEQ ID No. 34; SEQ ID No. 35; SEQ ID No. 39; SEQ ID No. 40; SEQ ID No. 43; SEQ ID No. 48; SEQ ID No. 49; SEQ ID No. 50; SEQ ID No. 52; SEQ ID No. 53; SEQ ID No. 54; SEQ ID No. 72; SEQ ID No. 77; SEQ ID No. 78; 25 SEQ ID No. 87; SEQ ID No. 90; SEQ ID No. 95; SEQ ID No. 108; SEQ ID No. 110; SEQ ID No. 122; SEQ ID No. 111; SEQ ID No. 123; SEQ ID No. 124; SEQ ID No. 127; SEQ ID No. 138; SEQ ID No. 144; SEQ ID No. 146; SEQ ID No. 153; SEQ ID No. 155; SEQ ID No. 164; SEQ ID No. 166; SEQ ID No. 175; SEQ ID No. 182; SEQ ID No. 184; SEQ ID No. 186; SEQ ID No. 187; SEQ ID No. 188; SEQ ID No. 202; SEQ ID No. 210; 30 SEQ ID No. 247; SEQ ID No. 258; SEQ ID No. 266; SEQ ID No. 267; SEQ ID No. 270; SEQ ID No. 273; SEQ ID No. 274; SEQ ID No. 295; SEQ ID No. 296; SEQ ID No. 305; SEQ ID No. 306; SEQ ID No. 309; SEQ ID No. 318; SEQ ID No. 319; SEQ ID No. 322; SEQ ID No. 326; SEQ ID No. 342; SEQ ID No. 357; SEQ ID No. 376; SEQ ID No. 379; SEQ ID No. 380; SEQ ID No. 388; SEQ ID No. 390; SEQ ID No. 400; SEQ ID No. 431;

SEQ ID No. 443;

SEQ ID No. 456;

SEQ ID No. 457;

SEQ ID No. 458; SEQ ID No. 464; SEQ ID No. 468; SEQ ID No. 470; SEQ ID No. 473; SEQ ID No. 489; SEQ ID No. 486; SEQ ID No. 497; SEQ ID No. 501; SEQ ID No. 503; SEQ ID No. 504; SEQ ID No. 508; SEQ ID No. 512; SEQ ID No. 521; SEQTD No. 522; SEQ ID No. 523; SEQ ID No. 524; SEQ ID No. 533; SEQ ID No. 535; SEQ ID No. 536; 5 SEQ ID No. 537; SEQ ID No. 538; SEQ ID No. 539; SEQ ID No. 540; SEQ ID No. 554; SEQ ID No. 563; SEQ ID No. 572; SEQ ID No. 579; SEQ ID No. 595; SEQ ID No. 603; SEQ ID No. 604; SEQ ID No. 606; SEQ ID No. 607; SEQ ID No. 615; SEQ ID No. 616; SEQ ID No. 622; SEQ ID No. 641; SEQ ID No. 642; SEQ ID No. 659; SEQ ID No. 668; SEQ ID No. 670; SEQ ID No. 693; SEQ ID No. 695; SEQ ID No. 696; SEQ ID No. 699; 10 SEQ ID No. 703; SEQ ID No. 704; SEQ ID No. 716; SEQ ID No. 726; SEQ ID No. 728; SEQ ID No. 739; SEQ ID No. 742; SEQ ID No. 747; SEQ ID No. 750; SEQ ID No. 751; SEQ ID No. 755; SEQ ID No. 757; SEQ ID No. 759; SEQ ID No. 761; SEQ ID No. 762; SEQ ID No. 763; SEQ ID No. 764; SEQ ID No. 773; SEQ ID No. 780; SEQ ID No. 781; SEQ ID No. 789; SEQ ID No. 800; SEQ ID No. 803; SEQ ID No. 804; SEQ ID No. 818; 15 SEQ ID No. 820; SEQ ID No. 822; SEQ ID No. 823; SEQ ID No. 824; SEQ ID No. 827; SEQ ID No. 828; SEQ ID No. 839; SEQ ID No. 849; SEQ ID No. 850; SEQ ID No. 851; SEQ ID No. 852; SEQ ID No. 855; SEQ ID No. 856; SEQ ID No. 857; SEQ ID No. 858; SEQ ID No. 859; SEQ ID No. 860; SEQ ID No. 861; SEQ ID No. 862; SEQ ID No. 863; SEQ ID No. 865; SEQ ID No. 868; SEQ ID No. 869; SEQ ID No. 870; SEQ ID No. 871; 20 SEQ ID No. 872; SEQ ID No. 873; SEQ ID No. 874; SEQ ID No. 875; SEQ ID No. 877; SEQ ID No. 878; SEQ ID No. 880; SEQ ID No. 882; SEQ ID No. 884; SEQ ID No. 886; SEQ ID No. 893; SEQ ID No. 901; SEQ ID No. 906; SEQ ID No. 910; SEQ ID No. 912; SEQ ID No. 915; SEQ ID No. 916; SEQ ID No. 917; SEQ ID No. 926; SEQ ID No. 929; SEQ ID No. 933; SEQ ID No. 965; SEQ ID No. 967; SEQ ID No. 968; SEQ ID No. 984; 25 SEQ ID No. 986; SEQ ID No. 989; SEQ ID No. 990; SEQ ID No. 996; SEQ ID No. 997; SEQ ID No. 1001; SEQ ID No. 1002; SEQ ID No. 1013; SEQ ID No. 1016; SEQ ID No. 1031; SEQ ID No. 1033; SEQ ID No. 1035; SEQ ID No. 1049; SEQ ID No. 1051; SEQ ID No. 1052; SEQ ID No. 1054; SEQ ID No. 1056; SEQ ID No. 1057; SEQ ID No. 1058; SEQ ID No. 1062; SEQ ID No. 1070; SEQ ID No. 1071; SEQ ID No. 1073 and one of their representative fragments. 30

In general, in the present invention, the functional group to which a polypeptide of the invention belongs, as well as its corresponding nucleotide sequence, may be determined either by comparative analogy with sequences already known, or by the use of standard techniques of biochemistry, of cytology combined with the techniques of genetic engineering such as immunoaffinity, localization by immunolabelling, differential extraction, measurement of enzymatic activity, study of the activity inducing or repressing expression or the study of expression in *E. coli*.

It is clearly understood, on the one hand, that, in the present invention, the nucleotide sequences (ORF) and the amino acid sequences (SEQ ID No. 2 to SEQ ID No. 1197) which are listed by functional group, are not exhaustive within the group considered. Moreover, it is also clearly understood that, in the present invention, a nucleotide sequence (ORF) or an amino acid sequence mentioned within a given functional group may also be part of another group taking into account, for example, the interrelationship between the groups listed. Accordingly, and as an example of this interrelationship, an exported and/or secreted polypeptide as well as its coding nucleotide sequence may also be involved in the *Chlamydia trachomatis* virulence process by modifying the defense mechanism of the infected host cell, or a transmembrane polypeptide or its coding nucleotide sequence is also part of the polypeptides or coding nucleotide sequences of the cellular envelope.

The subject of the present invention is also the nucleotide and/or polypeptide sequences according to the invention, characterized in that the said sequences are recorded on a medium, called recording medium, whose type and nature facilitate the reading, the analysis and the exploitation of the said sequences. These media may of course also contain other information extracted from the present invention, such as in particular the analogies with already known sequences, such as those mentioned in Table 1 of the present description, and/or may contain, in addition, information relating to the nucleotide and/or polypeptide sequences of other microorganisms so as to facilitate the comparative analysis and the exploitation of the results obtained.

Among these recording media, computer-readable media, such as magnetic, optical,
20 electrical and hybrid media such as, for example, floppy disks, CD-ROMs or recording cassettes, are
preferred in particular.

The invention also relates to nucleotide sequences which can be used as primer or probe, characterized in that the said sequences are chosen from the nucleotide sequences according to the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, as primer or probe, for the detection and/or amplification of nucleic acid sequences.

The nucleotide sequences according to the invention may thus be used to amplify nucleotide sequences, in particular by the PCR technique (polymerase chain reaction) (Erlich, 1989; Innis et al., 1990; Rolfs et al., 1991, and White et al., 1997).

These oligodeoxyribonucleotide or oligoribonucleotide primers correspond to representative nucleotide fragments, and are advantageously at least 8 nucleotides, preferably at least 12 nucleotides, 15 nucleotides and still more preferably at least 20 nucleotides long.

Other techniques for amplifying the target nucleic acid may be advantageously used as alternatives to PCR.

The nucleotide sequences of the invention, in particular the primers according to the

invention, may also be used in other methods for amplifying a target nucleic acid, such as:

- the TAS (Transcription-based Amplification System) technique described by Kwoh et al. in 1989;
- the 3SR (Self-Sustained Sequence Replication) technique described by Guatelli et al. in 1990;
- 5 the NASBA (Nucleic Acid Sequence Based Amplification) technique described by Kievitis et al. in 1991;
 - the SDA (Strand Displacement Amplification) technique (Walker et al., 1992);
 - the TMA (Transcription Mediated Amplification) technique.

The polynucleotides of the invention may also be used in techniques for amplifying or for modifying the nucleic acid serving as probe, such as:

- the LCR (Ligase Chain Reaction) technique described by Landegren et al. in 1988 and perfected by Barany et al. in 1991, which uses a thermostable ligase;
- the RCR (Repair Chain Reaction) technique described by Segev in 1992;
- the CPR (Cycling Probe Reaction) technique described by Duck et al. in 1990;
- the Q-beta-replicase amplification technique described by Miele et al. in 1983 and perfected in particular by Chu et al. in 1986, Lizardi et al. in 1988, and then by Burg et al. as well as by Stone et al. in 1996.

The invention also relates to the nucleotide sequences of fragments which can be obtained by amplification with the aid of at least one primer according to the invention. The present invention encompasses both hybridization probes and primers. In general, the complementary probes should be of the length sufficient to form a stable hybrid complex with the target sequences. Primers, while complementary to the target sequences need not form stable hybridization complexes with the target sequences alone. Rather, primers form stable complexes with the target sequences in the presence of polymerase to permit extension of the primer.

In the case where the target polynucleotide to be detected is possibly an RNA, for example an mRNA, it will be possible to use, prior to the use of an amplification reaction with the aid of at least one primer according to the invention or to the use of a method of detection with the aid of at least one probe of the invention, a reverse transcriptase-type enzyme so as to obtain a cDNA from the RNA contained in the biological sample. The cDNA obtained will then serve as target for the primer(s) or the probe(s) used in the amplification or detection method according to the invention.

The detection probe will be chosen so that it hybridizes with the target sequence or the amplicon generated from the target sequence. Such a detection probe will advantageously have as sequence a sequence of at least 12 nucleotides, 15 nucleotides, in particular of at least 20 nucleotides, and preferably at least 100 nucleotides.

The invention also comprises the nucleotide sequences which can be used as probe or

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primer according to the invention, characterized in that they are labelled with a radioactive compound or with a nonradioactive compound.

The nonlabelled nucleotide sequences may be used directly as probes or primers; however, the sequences are generally labelled with a radioactive element (³²P, ³⁵S, ³H, ¹²⁵I) or with a nonradioactive molecule (biotin, acetylaminofluorene, digoxigenin, 5-bromo-deoxyuridine, fluorescein) so as to obtain probes which can be used in numerous applications.

Examples of nonradioactive labelling of nucleotide sequences are described, for example, in French patent No. 78,10975 or by Urdea et al. or by Sanchez-Pescador et al. in 1988.

In the latter case, one of the labelling methods described in patents FR-2 422 956 and FR-2 518 755 may also be used.

The invention also relates to the nucleotide sequences of fragments which can be obtained by hybridization with the aid of at least one probe according to the invention.

The hybridization technique may be performed in various ways (Matthews et al., 1988). The most common method consists in immobilizing the nucleic acid extracted from C. trachomatis cells on a support (such as nitrocellulose, nylon, polystyrene) and in incubating, under well-defined conditions, the target nucleic acid immobilized with the probe. After hybridization, the excess probe is removed and the hybrid molecules formed are detected by the appropriate method (measurement of the radioactivity, of the fluorescence or of the enzymatic activity linked to the probe).

The invention also comprises the nucleotide sequences according to the invention, characterized in that they are covalently or noncovalently immobilized on a support.

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According to another advantageous embodiment of the nucleic sequences according to the invention, the latter may be used immobilized on a support and may thus serve to capture, through specific hybridization, the target nucleic acid obtained from the biological sample to be tested. If necessary, the solid support is separated from the sample and the hybridization complex formed between the so-called capture probe and the target nucleic acid is then detected by means of a second probe, called detection probe, labelled with an easily detectable element.

The nucleotide sequences according to the invention may also be used in new analytical systems, DNA chips, which allow sequencing, the study of mutations and of the expression of genes, and which are currently of interest given their very small size and their high capacity in terms of number of analyses.

The principle of the operation of these chips is based on molecular probes, most often oligonucleotides, which are attached onto a miniaturized surface, generally of the order of a few square centimetres. During an analysis, a sample containing fragments of a target nucleic acid to be analysed, for example DNA or RNA labelled, for example, after amplification, is deposited onto the

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DNA chip in which the support has been coated beforehand with probes. Bringing the labelled target sequences into contact with the probes leads to the formation, through hybridization, of a duplex according to the rule of pairing defined by J.D. Watson and F. Crick. After a washing step, analysis of the surface of the chip allows the effective hybridizations to be located by means of the signals emitted by the labels tagging the target. A hybridization fingerprint results from this analysis which, by appropriate computer processing, will make it possible to determine information such as the presence of specific fragments in the sample, the determination of sequences and the presence of mutations.

The chip consists of a multitude of molecular probes, precisely organized or arrayed on a solid support whose surface is miniaturized. It is at the centre of a system where other elements (imaging system, microcomputer) allow the acquisition and interpretation of a hybridization fingerprint.

The hybridization supports are provided in the form of flat or porous surfaces (pierced with wells) composed of various materials. The choice of a support is determined by its physicochemical properties, or more precisely, by the relationship between the latter and the conditions under which the support will be placed during the synthesis or the attachment of the probes or during the use of the chip. It is therefore necessary, before considering the use of a particular support (R.S. Matson et al., 1994), to consider characteristics such as its stability to pH, its physical strength, its reactivity and its chemical stability as well as its capacity to nonspecifically bind nucleic acids. Materials such as glass, silicon and polymers are commonly used. Their surface is, in a first step, called «functionalization», made reactive towards the groups which it is desired to attach thereon. After the functionalization, so-called spacer molecules are grafted onto the activated surface. Used as intermediates between the surface and the probe, these molecules of variable size render unimportant the surface properties of the supports, which often prove to be problematic for the synthesis or the attachment of the probes and for the hybridization.

Among the hybridization supports, there may be mentioned glass which is used, for example, in the method of in situ synthesis of oligonucleotides by photochemical addressing developed by the company Affymetrix (E.L. Sheldon, 1993), the glass surface being activated by silane. Genosensor Consortium (P. Mérel, 1994) also uses glass slides carrying wells 3 mm apart, this support being activated with epoxysilane.

Polymers or silicon may also be mentioned among these hybridization supports. For example, the Andrein Mirzabekov team has developed a chip consisting of polyacrylamide squares polymerized on a silanized glass surface (G. Yershov et al., 1996). Several teams use silicon, in particular the IFOS laboratory of Ecole Centrale of Lyon which uses a silicon semiconductor substrate which is p-doped by introducing it into its crystalline structure atoms whose valency is different from

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that of silicon. Various types of metals, in particular gold and platinum, may also be used as support (Genosensor Consortium (K. Beattie et al., 1993)).

The probes according to the invention may be synthesized directly in situ on the supports of the DNA chips. This in situ synthesis may be carried out by photochemical addressing (developed by the company Affymax (Amsterdam, Holland) and exploited industrially by its subsidiary Affymetrix (United States)) or based on the VLSIPS (very large scale immobilized polymer synthesis) technology (S.P.A. Fodor et al., 1991) which is based on a method of photochemically directed combinatory synthesis and the principle of which combines solid-phase chemistry, the use of photolabile protecting groups and photolithography.

The probes according to the invention may be attached to the DNA chips in various ways such as electrochemical addressing, automated addressing or the use of probe printers (T. Livache et al., 1994; G. Yershov et al., 1996; J. Derisi et al., 1996, and S. Borman, 1996).

The revealing of the hybridization between the probes of the invention, deposited or synthesized in situ on the supports of the DNA chips, and the sample to be analysed, may be determined, for example, by measurement of fluorescent signals, by radioactive counting or by electronic detection.

The use of fluorescent molecules such as fluorescein constitutes the most common method of labelling the samples. It allows direct or indirect revealing of the hybridization and allows the use of various fluorochromes.

Affymetrix currently provides an apparatus or a scanner designed to read its Gene Chip? chips. It makes it possible to detect the hybridizations by scanning the surface of the chip in confocal microscopy (R.J. Lipshutz et al., 1995). Other methods of detecting fluorescent signals have been tested: coupling of an epifluorescence microscope and a CCD camera (G. Yershov et al., 1996), the use of an optical fibre collecting system (E.L. Sheldon, 1993). A conventional method consists in carrying out an end labelling, with phosphorus 32, of the target sequences, by means of an appropriate apparatus, the Phosphorimager (marketed by Molecular Dynamics). The electronic detection is based on the principle that the hybridization of two nucleic acid molecules is accompanied by physical phenomena which can be quantified under certain conditions (system developed by Ecole Centrale of Lyon and called GEN-FET (GEN field effect transistor)). Genosensor Consortium and the company Beckman Instruments who are developing an electronic chip or Permittivity Chips? may also be mentioned (K. Beattie et al., 1993).

The nucleotide sequences according to the invention may thus be used in DNA chips to carry out the analysis of mutations. This analysis is based on the production of chips capable of analysing each base of a nucleotide sequence according to the invention. It is possible, in particular to this end, to use the microsequencing techniques on a DNA chip. The mutations are detected by

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extending immobilized primers which hybridize to the template of sequences analysed, just at the position adjacent to that of the mutated nucleotide to be detected. A single-stranded template, RNA or DNA, of the sequences to be analysed will be advantageously prepared according to conventional methods, from products amplified according to PCR-type techniques. The templates of single-5 stranded DNA, or of RNA thus obtained are then deposited on the DNA chip, under conditions allowing their specific hybridization to the immobilized primers. A thermostable polymerase, for example Tth or T7 DNA polymerase, specifically extends the 3' end of the immobilized primer with a labelled nucleotide analogue complementary to the nucleotide at the position of the variable site. For example a thermal cycling is performed in the presence of fluorescent dideoxyribonucleotides. 10 The experimental conditions will be adapted in particular to the chips used, to the immobilized primers, to the polymerases used and to the labelling system chosen. One advantage of microsequencing, compared with techniques based on the hybridization of probes, is that it makes it possible to identify all the variable nucleotides with optimal discrimination under homogeneous reaction conditions; used on DNA chips, it allows optimal resolution and specificity for the routine and industrial detection of mutations in multiplex.

The nucleotide sequences according to the invention may also be used in DNA chips to carry out the analysis of the expression of the Chlamydia trachomatis genes. This analysis of the expression of Chlamydia trachomatis genes is based on the use of chips where probes of the invention, chosen for their specificity to characterize a given gene, are present (D.J. Lockhart et al., 20 1996; D.D. Shoemaker et al., 1996). For the methods of analysis of gene expression using the DNA chips, reference may, for example, be made to the methods described by D.J. Lockhart et al. (1996) and Sosnowsky et al. (1997) for the synthesis of probes in situ or for the addressing and the attachment of previously synthesized probes. The target sequences to be analysed are labelled and in general fragmented into sequences of about 50 to 100 nucleotides before being hybridized onto the 25 chip. After washing as described, for example, by D.J. Lockhart et al. (1996) and application of different electric fields (Sosnowsky et al., 1997), the labelled compounds are detected and quantified, the hybridizations being carried out at least in duplicate. Comparative analyses of the signal intensities obtained with respect to the same probe for different samples and/or for different probes with the same sample, determine the differential expression of RNA or of DNA derived from the sample.

The nucleotide sequences according to the invention may, in addition, be used in DNA chips where other nucleotide probes specific for other microorganisms are also present, and may allow the carrying out of a serial test allowing rapid identification of the presence of a microorganism in a sample.

to the invention, characterized in that they are immobilized on a support of a DNA chip.

The DNA chips, characterized in that they contain at least one nucleotide sequence according to the invention, immobilized on the support of the said chip, also form part of the invention.

The said chips will preferably contain several probes or nucleotide sequences of the invention of different length and/or corresponding to different genes so as to identify, with greater certainty, the specificity of the target sequences or the desired mutation in the sample to be analysed.

Accordingly, the analyses carried out by means of primers and/or probes according to the invention, immobilized on supports such as DNA chips, will make it possible, for example, to identify, in samples, mutations linked to variations such as intraspecies variations. These variations may be correlated or associated with pathologies specific to the variant identified and will make it possible to select the appropriate treatment.

The invention thus comprises a DNA chip according to the invention, characterized in that it contains, in addition, at least one nucleotide sequence of a microorganism different from Chlamydia trachomatis, immobilized on the support of the said chip; preferably, the different microorganism will be chosen from an associated microorganism, a bacterium of the Chlamydia family, and a variant of the species Chlamydia trachomatis.

Another subject of the present invention is a vector for the cloning and/or the expression of a sequence, characterized in that it contains a nucleotide sequence according to the invention.

Among the said vectors according to the invention, the vectors containing a nucleotide sequence encoding a polypeptide of the cellular, preferably outer, envelope of *Chlamydia trachomatis* or one of its representative fragments, are preferred.

In a specific embodiment, the vectors contain a nucleotide sequence encoding a Chlamydia trachomatis secreted polypeptide or one of its representative fragments or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its representative fragments, a polypeptide involved in lipopolysaccharide (LPS) biosynthesis, a Type III or non-Type III secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall anchored surface polypeptide, a polypeptide not found in Chlamydia pneumoniae, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids or of fatty acids of Chlamydia trachomatis or one of their representative fragments, or a polypeptide specific to Chlamydiae, are also preferred.

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According to the invention, the vectors comprise the elements necessary to allow the expression and/or the secretion of the said nucleotide sequences in a given host cell, and also form part of the invention.

The vector should, in this case, comprise a promoter, signals for initiation and for termination of translation, as well as appropriate regions for regulation of transcription. It should be capable of being stably maintained in the host cell and may optionally possess particular signals specifying the secretion of the translated protein. These different elements are chosen according to the host cell used. To this effect, the nucleotide sequences according to the invention may be inserted into autonomously-replicating vectors within the chosen host, or integrative vectors in the chosen host.

10 Any of the standard methods known to those skilled in the art for the insertion of

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DNA fragments into a vector may be used to construct expression vectors containing a chimeric gene consisting of appropriate transcriptional/translational control signals and the protein coding sequences. These methods may include in vitro recombinant DNA and synthetic techniques and in

vivo recombinants (genetic recombination).

Expression of a polypeptide, peptide or derivative, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1 or ORFs contained within SEQ ID No. 1 may be regulated by a second nucleic acid sequence so that the protein or peptide is expressed in a host transformed with the recombinant DNA molecule. For example, expression of a protein or peptide may be controlled by any promoter/enhancer element known in the art. Promoters which may be used to 20 control expression include, but are not limited to, the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3' long terminal repeat of Rous sarcoma virus (Yamamoto, et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42); prokaryotic expression vectors 25 such as the β-lactamase promoter (Villa-Kamaroff, et al., 1978, Proc. Natl. Acad. Sci. U.S.A. 75:3727-3731), or the tac promoter (DeBoer, et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:21-25); see also "Useful proteins from recombinant bacteria" in Scientific American, 1980, 242:74-94; plant expression vectors comprising the nopaline synthetase promoter region (Herrera-Estrella et al., 1983, Nature 303:209-213) or the cauliflower mosaic virus 35S RNA promoter (Gardner, et al., 1981, Nucl. 30 Acids Res. 2:2871), and the promoter of the photosynthetic enzyme ribulose biphosphate carboxylase (Herrera-Estrella et al., 1984, Nature 310:115-120); promoter elements from yeast or other fungi such as the Gal 4 promoter, the ADC (alcohol dehydrogenase) promoter, PGK (phosphoglycerol kinase) promoter, alkaline phosphatase promoter, and the following animal transcriptional control regions, which exhibit tissue specificity and have been utilized in transgenic animals: elastase I gene control region which is active in pancreatic acinar cells (Swift et al., 1984, Cell 38:639-646; Ornitz et al.,

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1986, Cold Spring Harbor Symp. Quant. Biol. 50:399-409; MacDonald, 1987, Hepatology 7:425-515); insulin gene control region which is active in pancreatic beta cells (Hanahan, 1985, Nature 315:115-122), immunoglobulin gene control region which is active in lymphoid cells (Grosschedl et al., 1984, Cell 38:647-658; Adames et al., 1985, Nature 318:533-538; Alexander et al., 1987, Mol. 5 Cell. Biol. 7:1436-1444), mouse mammary tumor virus control region which is active in testicular, breast, lymphoid and mast cells (Leder et al., 1986, Cell 45:485-495), albumin gene control region which is active in liver (Pinkert et al., 1987, Genes and Devel. 1:268-276), alpha-fetoprotein gene control region which is active in liver (Krumlauf et al., 1985, Mol. Cell. Biol. 5:1639-1648; Hammer et al., 1987, Science 235:53-58; alpha 1-antitrypsin gene control region which is active in the liver 10 (Kelsey et al., 1987, Genes and Devel. 1:161-171), beta-globin gene control region which is active in myeloid cells (Mogram et al., 1985, Nature 315:338-340; Kollias et al., 1986, Cell 46:89-94; myelin basic protein gene control region which is active in oligodendrocyte cells in the brain (Readhead et al., 1987, Cell 48:703-712); myosin light chain-2 gene control region which is active in skeletal muscle (Sani, 1985, Nature 314:283-286), and gonadotropic releasing hormone gene control region which is active in the hypothalamus (Mason et al., 1986, Science 234:1372-1378).

The vectors according to the invention are, for example, vectors of plasmid or viral origin. In a specific embodiment, a vector is used that comprises a promoter operably linked to a protein or peptide-encoding nucleic acid sequence in SEQ ID No. 1, or ORFs contained within SEQ ID No. 1, one or more origins of replication, and, optionally, one or more selectable markers (e.g., an antibiotic resistance gene). Expression vectors comprise regulatory sequences that control gene expression, including gene expression in a desired host cell. Preferred vectors for the expression of the polypeptides of the invention include the pET-type plasmid vectors (Promega) or pBAD plasmid vectors (Invitrogen). Furthermore, the vectors according to the invention are useful for transforming host cells so as to clone or express the nucleotide sequences of the invention.

Expression can also be achieved using targeted homologous recombination to activate Chlamydia trachomatis genes present in the cloned genomic DNA. A heterologous regulatory element may be inserted into a stable cell line or cloned microorganism, such that it is operatively linked with an endogenous Chlamydia trachomatis gene present in the cloned genome, using techniques, such as targeted homologous recombination, which are well known to those of skill in the art (See, e.g., Chappel, U.S. Patent No. 4,215,051 and Skoultchi, WO 91/06667 each of which is incorporated herein in its entirety).

Expression vector/host cell systems containing inserts of polynucleotide sequences in SEQ ID No. 1 or ORFs within SEQ ID No. 1, which encode polypeptides, peptides or derivatives, or analogs thereof, can be identified by three general approaches: (a) nucleic acid hybridization, (b) presence or absence of "marker" gene functions, and (c) expression of inserted sequences. In the first WO 99/28475 PCT/IB98/01939

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approach, the presence of a polynucleotide sequence inserted in an expression vector can be detected by nucleic acid hybridization using probes comprising sequences that are homologous to an inserted polynucleotide sequence. In the second approach, the recombinant vector/host system can be identified and selected based upon the presence or absence of certain "marker" gene functions (e.g., thymidine kinase activity, resistance to antibiotics, transformation phenotype, occlusion body formation in baculovirus, etc.) caused by the insertion of a polynucleotide sequence in the vector. For example, if the polynucleotide sequence in SEQ ID No. 1 or ORFs within SEQ ID No. 1 is inserted within the marker gene sequence of the vector, recombinants containing the insert can be identified by the absence of the marker gene function. In the third approach, recombinant expression vectors can be identified by assaying the product of the polynucleotide sequence expressed by the recombinant. Such assays can be based, for example, on the physical or functional properties of the expressed polypeptide in in vitro assay systems, e.g., binding with antibody, promotion of cell proliferation.

Once a particular recombinant DNA molecule is identified and isolated, several methods known in the art may be used to propagate it. The clones identified may be introduced into an appropriate host cell by standard methods, such as for example lipofection, electroporation, and heat shock. Once a suitable host system and growth conditions are established, recombinant expression vectors can be propagated and prepared in quantity.

The invention also encompasses the host cells transformed by a vector according to the invention. These cells may be obtained by introducing into host cells a nucleotide sequence inserted into a vector as defined above, and then culturing the said cells under conditions allowing the replication and/or the expression of the transfected nucleotide sequence.

The host cell may be chosen from eukaryotic or prokaryotic systems, such as for example bacterial cells (Olins and Lee, 1993), but also yeast cells (Buckholz, 1993), as well as animal cells, in particular cultures of mammalian cells (Edwards and Aruffo, 1993), and in particular Chinese hamster ovary (CHO) cells, but also insect cells in which methods using baculoviruses for example may be used (Luckow, 1993).

Furthermore, a host cell strain may be chosen which modulates the expression of the inserted sequences, or modifies and processes the gene product in the specific fashion desired. Expression from certain promoters can be elevated in the presence of certain inducers; thus, expression of the genetically engineered polypeptide may be controlled. Furthermore, different host cells have characteristic and specific mechanisms for the translational and post-translational processing and modification (e.g., glycosylation, phosphorylation) of proteins. Appropriate cell lines or host systems can be chosen to ensure the desired modification and processing of the foreign protein expressed. For example, expression in a bacterial system can be used to produce an unglycosylated core protein product. Expression in yeast will produce a glycosylated product. Expression in

mammalian cells can be used to ensure "native" glycosylation of a heterologous protein. Furthermore, different vector/host expression systems may effect processing reactions to different extents.

A preferred host cell for the expression of the proteins of the invention consists of prokaryotic cells, such as Gram negative bacteria.

A further preferred host cell according to the invention is a bacterium belonging to the *Chlamydia* family, more preferably belonging to the species *Chlamydia trachomatis* or chosen from a microorganism associated with the species *Chlamydia trachomatis*.

In other specific embodiments, the polypeptides, peptides or derivatives, or analogs thereof may be expressed as a fusion, or chimeric protein product (comprising the protein, fragment, analog, or derivative joined via a peptide bond to a heterologous protein sequence (of a different protein)). Such a chimeric product can be made by ligating the appropriate nucleic acid sequences encoding the desired amino acid sequences to each other by methods known in the art, in the proper coding frame, and expressing the chimeric product by methods commonly known in the art. Alternatively, such a chimeric product may be made by protein synthetic techniques, e.g., by use of a peptide synthesizer.

Genomic sequences can be cloned and expressed as translational gene products (<u>i.e.</u>, peptides, polypeptides, and proteins) or transcriptional gene products (<u>i.e.</u>, antisense and ribozymes).

The invention further relates to the intracellular production of an antisense nucleic acid sequence of SEQ ID No. 1 by transcription from an exogenous sequence. For example, a vector can be introduced *in vivo* such that it is taken up by a cell, within which cell the vector or a portion thereof is transcribed, producing an antisense nucleic acid (RNA) of the invention. Such a vector would contain a sequence encoding an antisense nucleic acid. Such a vector can remain episomal or become chromosomally integrated, as long as it can be transcribed to produce the desired antisense RNA. Such vectors can be constructed by recombinant DNA technology methods standard in the art.

Vectors can be plasmid, viral, or others known in the art, used for replication and expression in mammalian cells. Expression of the sequence encoding the antisense RNA can be by any promoter known in the art to act in mammalian, preferably human, cells. Such promoters can be inducible or constitutive. Such promoters include but are not limited to: the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3N long terminal repeat of Rous sarcoma virus (Yamamoto et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42), etc.

In a specific embodiment, the antisense oligonucleotide comprises catalytic RNA, or a ribozyme (see, e.g., PCT International Publication WO 90/11364, published October 4, 1990; Sarver et al., 1990, Science 247:1222-1225). In another embodiment, the oligonucleotide is a 2N-0-

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methylribonucleotide (Inoue et al., 1987, Nucl. Acids Res. <u>15</u>:6131-6148), or a chimeric RNA-DNA analog (Inoue et al., 1987, FEBS Lett. <u>215</u>:327-330).

In another embodiment, the antisense nucleic acids of the invention comprise a sequence complementary to at least a portion of an RNA transcript of a polynucleotide sequence in SEQ ID No. 1. However, absolute complementarity, although preferred, is not required. A sequence "complementary to at least a portion of an RNA," as referred to herein, means a sequence having sufficient complementarity to be able to hybridize with the RNA, forming a stable duplex; in the case of double-stranded antisense nucleic acid sequence, a single strand of the duplex DNA may thus be tested, or triplex formation may be assayed. The ability to hybridize will depend on both the degree of complementarity and the length of the antisense nucleic acid. Generally, the longer the hybridizing nucleic acid, the more base mismatches with an RNA transcribed from SEQ ID No. 1 may contain and still form a stable duplex (or triplex, as the case may be). One skilled in the art can ascertain a tolerable degree of mismatch by use of standard procedures to determine the melting point of the hybridized complex.

The invention also relates to the animals, except humans, comprising one of the above-described transformed cells according to the invention.

The production of transgenic animals according to the invention overexpressing one or more of the *Chlamydia trachomatis* genes will be preferably carried out on rats, mice or rabbits according to methods well known to persons skilled in the art such as viral or nonviral transfections.

The transgenic animals overexpressing one or more of the said genes may be obtained by transfection of multiple copies of the said genes under the control of a powerful promoter of a ubiquitous nature, or which is selective for one type of tissue. The transgenic animals may also be obtained by homologous recombination on embryonic stem cells, transfer of these stem cells to embryos, selection of the chimeras affected at the level of the reproductive lines, and growth of the said chimeras.

The transformed cells as well as the transgenic animals according to the invention can be used in methods of preparing the recombinant polypeptide.

It is now possible to produce recombinant polypeptides in a relatively large quantity by genetic engineering using the cells transformed with expression vectors according to the invention or using transgenic animals according to the invention.

The methods of preparing a polypeptide of the invention in recombinant form, characterized in that they use a vector and/or a cell transformed with a vector according to the invention and/or a transgenic animal comprising one of the said transformed cells according to the invention, are themselves included in the present invention.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or

a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a polypeptide of the cellular envelope of Chlamydia trachomatis or one of its representative fragments, more preferably encoding a polypeptide of the outer cellular envelope of Chlamydia trachomatis or one of its fragment, are preferred.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a Chlamydia trachomatis secreted polypeptide or one of its representative fragments, or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its 10 representative fragments, a polypeptide involved in lipopolysaccharide biosynthesis, a Type III or other secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall anchored surface polypeptide, a polypeptide not found in Chlamydia pneumoniae, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved 15 in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids or of fatty acids of Chlamydia trachomatis or one of their representative fragments, or a polypeptide specific to Chlamydiae, are also preferred.

The recombinant polypeptides obtained as indicated above may be provided either in 20 glycosylated or nonglycosylated form and may or may not have the natural tertiary structure.

A preferred variant consists in producing a recombinant polypeptide fused to a «carrier» protein (chimeric protein). The advantage of this system is that it allows a stabilization and a reduction in proteolysis of the recombinant product, an increase in solubility during renaturation in vitro and/or a simplification of purification when the fusion partner has affinity for a specific 25 ligand.

More particularly, the invention relates to a method of preparing a polypeptide of the invention comprising the following steps:

- a) culture of the transformed cells under conditions allowing the expression of a recombinant polypeptide having a nucleic acid sequence according to the invention;
- 30 b) where appropriate, recovery of the said recombinant polypeptide.

When the method of preparing a polypeptide of the invention uses a transgenic animal according to the invention, the recombinant polypeptide is then extracted from the said animal.

The subject of the invention is also a polypeptide capable of being obtained by a method of the invention as described above.

35 The invention also comprises a method of preparing a synthetic polypeptide,

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characterized in that it uses an amino acid sequence of polypeptides according to the invention.

The invention also relates to a synthetic polypeptide obtained by a method according to the invention.

Polypeptides according to the invention may also be prepared by conventional 5 techniques in the field of peptide synthesis under conditions suitable to produce the polypeptides encoded by the polynucleotide of the invention. This synthesis may be carried out in and recovered from a homogeneous solution or on a solid phase.

For example, the synthesis technique in a homogeneous solution described by Houbenweyl in 1974 may be used.

This method of synthesis consists in successively condensing, in pairs, the successive amino acids in the required order, or in condensing amino acids and fragments previously formed and already containing several amino acids in the appropriate order, or alternatively several fragments thus previously prepared, it being understood that care will have been taken to protect beforehand all the reactive functional groups carried by these amino acids or fragments, with the exception of the amine functional groups of one and the carboxyl functional groups of the other or vice versa, which should normally take part in the formation of the peptide bonds, in particular after activation of the carboxyl functional group, according to methods well known in peptide synthesis.

According to another preferred technique of the invention, the one described by Merrifield is used.

To manufacture a peptide chain according to the Merrifield method, a highly porous polymer resin is used, onto which the first C-terminal amino acid of the chain is attached. This amino acid is attached onto a resin via its carboxyl group and its amine functional group is protected. The amino acids which will constitute the peptide chain are thus attached, one after another, onto the amine group, each time deprotected beforehand, of the portion of the peptide chain already formed, 25 and which is attached to the resin. When the entire peptide chain desired is formed, the protecting groups are removed from the various amino acids constituting the peptide chain and the peptide is detached from the resin with the aid of an acid.

The invention relates, in addition, to hybrid (fusion) polypeptides having at least one polypeptide or one of its representative fragments according to the invention, and a sequence of a polypeptide capable of eliciting an immune response in humans or animals.

Advantageously, the antigenic determinant is such that it is capable of eliciting a humoral and/or cellular response.

An antigenic determinant may be identified by screening expression libraries of the Chlamydia trachomatis genome with antibodies contained in the serum of patients infected with a bacterium belonging to the species Chlamydia trachomatis. An antigenic determinant may comprise a

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polypeptide or one of its fragments according to the invention, in glycosylated form, used in order to obtain immunogenic compositions capable of inducing the synthesis of antibodies directed against multiple epitopes. The said polypeptides or their glycosylated fragments also form part of the invention.

These hybrid molecules may consist, in part, of a carrier molecule for polypeptides or for their representative fragments according to the invention, combined with a portion which may be immunogenic, in particular an epitope of the diphtheria toxin, the tetanus toxin, a hepatitis B virus surface antigen (patent FR 79 21811), the poliomyelitis virus VP1 antigen or any other viral or bacterial toxin or antigen.

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The methods of synthesizing the hybrid molecules include the methods used in genetic engineering to construct hybrid nucleotide sequences encoding the desired polypeptide sequences. Reference may be advantageously made, for example, to the technique for producing genes encoding fusion proteins described by Minton in 1984.

The said hybrid nucleotide sequences encoding a hybrid polypeptide as well as the hybrid polypeptides according to the invention, characterized in that they are recombinant polypeptides obtained by the expression of the said hybrid nucleotide sequences, also form part of the invention.

The invention also comprises the vectors characterized in that they contain one of the said hybrid nucleotide sequences. The host cells transformed by the said vectors, the transgenic animals comprising one of the said transformed cells as well as the methods of preparing recombinant polypeptides using the said vectors, the said transformed cells and/or the said transgenic animals of course also form part of the invention.

The polypeptides according to the invention, the antibodies according to the invention described below and the nucleotide sequences according to the invention may advantageously be used in in vitro and/or in vivo methods for the detection and/or the identification of bacteria belonging to the species Chlamydia trachomatis, in a biological sample (biological tissue or fluid) which is likely to contain them. These methods, depending on the specificity of the polypeptides, of the antibodies and of the nucleotide sequences according to the invention which will be used, may in particular detect and/or identify the bacterial variants belonging to the species Chlamydia trachomatis as well as the associated microorganisms capable of being detected by the polypeptides, the antibodies and the nucleotide sequences according to the invention which will be chosen. It may, for example, be advantageous to choose a polypeptide, an antibody or a nucleotide sequence according to the invention, which is capable of detecting any bacterium of the Chlamydia family by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the family or, on the contrary, it will be most particularly advantageous to target a variant of the

species Chlamydia trachomatis, which is responsible, for example, for the induction or the worsening of pathologies specific to the targeted variant, by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the said variant.

The polypeptides according to the invention may advantageously be used in a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, in a biological sample (biological tissue or fluid) which is likely to contain them, characterized in that it comprises the following steps:

- a) bringing this biological sample into contact with a polypeptide or one of its representative fragments according to the invention (under conditions allowing an immunological reaction between
 the said polypeptide and the antibodies which may be present in the biological sample);
 - b) detecting the antigen-antibody complexes which may be formed.

Preferably, the biological sample consists of a fluid, for example a human or animal serum, blood or biopsies.

Any conventional procedure may be used to carry out such a detection of the antigenantibody complexes which may be formed.

By way of example, a preferred method uses immunoenzymatic procedures based on the ELISA technique, immunofluorescence procedures or radioimmunological procedures (RIA), and the like.

Accordingly, the invention also relates to the polypeptides according to the invention, labelled with the aid of a suitable label such as a label of the enzymatic, fluorescent or radioactive type.

Such methods comprise, for example, the following steps:

- deposition of defined quantities of a polypeptide composition according to the invention into the wells of a microtitre plate,
- 25 introduction, into the said wells, of increasing dilutions of serum, or of a different biological sample as defined above, which has to be analysed.
 - incubation of the microplate,
- introduction, into the wells of the microtitre plate, of labelled antibodies directed against human or animal immunoglobulins, these antibodies having been labelled with the aid of an enzyme selected from those which are capable of hydrolyzing a substrate, thereby modifying the absorption of the radiation of the latter, at least at a defined wavelength, for example at 550 nm,
 - detection, by comparison with a control, of the quantity of substrate hydrolyzed.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a polypeptide according to the invention,
- where appropriate, the reagents for constituting the medium appropriate for the immunological or specific reaction,
- the reagents allowing the detection of the antigen-antibody complexes produced by the immunological reaction between the polypeptide(s) of the invention and the antibodies which may be present in the biological sample, it being possible for these reagents also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the polypeptide according to the invention is not labelled,
- where appropriate, a reference biological sample (negative control) free of antibodies recognized by a polypeptide according to the invention,
 - where appropriate, a reference biological sample (positive control) containing a predetermined quantity of antibodies recognized by a polypeptide according to the invention.

According to the invention, the polypeptides, peptides, fusion proteins or other derivatives, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1, may be used as an immunogen to generate antibodies which immunospecifically bind such an immunogen. Such antibodies may include, but are not limited to, polyclonal and monoclonal antibodies, humanized or chimeric antibodies, single chain antibodies, Fab fragments, F(ab)₂ fragments, fragments produced by a Fab expression library, anti-idiotypic (anti-Id) antibodies, and epitope-binding fragments of any of the above. In a specific embodiment, the antibody to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1 is a bispecific antibody (see generally, e.g. Fanger and Drakeman, 1995, Drug News and Perspectives 8: 133-137). Such a bispecific antibody is genetically engineered to recognize both (1) an epitope and (2) one of a variety of "trigger" molecules, e.g. Fc receptors on myeloid cells, and CD3 and CD2 on T cells, that have been identified as being able to cause a cytotoxic T-cell to destroy a particular target. Such bispecific antibodies can be prepared either by chemical conjugation, hybridoma, or recombinant molecular biology techniques known to the skilled artisan.

Various procedures known in the art may be used for the production of polyclonal antibodies to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1. For the production of antibody, various host animals can be immunized by injection with a polypeptide, or peptide or other derivative, or analog thereof, including but not limited to rabbits, mice, rats, etc. Various adjuvants, depending on the host species, may be used to increase the immunological response, including but not limited to StimulonTM QS-21 (Aquila Biopharmaceuticals, Inc., Framingham, MA), MPLTM (3-O-deacylated monophosphoryl lipid A; RIBI ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA), Freund's (complete and incomplete), mineral gels such as aluminum hydroxide,

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surface active substances such as lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, keyhole limpet hemocyanins, dinitrophenol, BCG (bacille Calmette-Guerin), and corynebacterium parvum. Alternatively, polyclonal antibodies may be prepared by purifying, on an affinity column onto which a polypeptide according to the invention has been previously attached, the antibodies contained in the serum of patients infected with a bacterium belonging to the species *Chlamydia trachomatis*.

For preparation of monoclonal antibodies directed toward a polypeptide, peptide or other derivative, or analog, any technique which provides for the production of antibody molecules by continuous cell lines in culture may be used. For example, the hybridoma technique originally developed by Kohler and Milstein (1975, Nature 256:495-497), as well as the trioma technique, the human B-cell hybridoma technique (Kozbor et al., 1983, Immunology Today 4:72), and the EBVhybridoma technique to produce human monoclonal antibodies (Cole et al., 1985, in Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, Inc., pp. 77-96). In an additional embodiment of the invention, monoclonal antibodies can be produced in germ-free animals utilizing technology 15 described in PCT/US90/02545. In another embodiment of the invention, transgenic non-human animals can be used for the production of human antibodies utilizing technology described in WO 98/24893 and WO 96/33735. According to the invention, human antibodies may be used and can be obtained by using human hybridomas (Cote et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:2026-2030) or by transforming human B cells with EBV virus in vitro (Cole et al., 1985, in Monoclonal 20 Antibodies and Cancer Therapy, Alan R. Liss, pp. 77-96). In fact, according to the invention, techniques developed for the production of «chimeric antibodies», (Morrison et al., 1984, PROC. NATL. ACAD. SCI. U.S.A. 81:6851-6855; Neuberger et al., 1984, Nature 312:604-608; Takeda et al., 1985, Nature 314:452-454) by splicing the genes from a mouse antibody molecule specific for a polypeptide, peptide or other derivative, or analog together with genes from a human antibody molecule of appropriate biological activity can be used; such antibodies are within the scope of this invention.

According to the invention, techniques described for the production of single chain antibodies (U.S. Patent 4,946,778) can be adapted to produce polypeptide or peptide-specific single chain antibodies. An additional embodiment of the invention utilizes the techniques described for the construction of Fab expression libraries (Huse et al., 1989, Science 246:1275-1281) to allow rapid and easy identification of monoclonal Fab fragments with the desired specificity for polypeptides, derivatives, or analogs.

Antibody fragments which contain the idiotype of the molecule can be generated by known techniques. For example, such fragments include but are not limited to: the F(ab')₂ fragment which can be produced by pepsin digestion of the antibody molecule; the Fab' fragments which can be

generated by reducing the disulfide bridges of the F(ab')₂ fragment, the Fab fragments which can be generated by treating the antibody molecule with papain and a reducing agent, and Fv fragments.

In addition, techniques have been developed for the production of chimerized (See, e.g., Boss, M. et al., U.S. Patent No. 4,816,397; and Cabilly, S. et al., U.S. Patent No. 5,585,089 each of which is incorporated herein by reference in its entirety) humanized antibodies (See, e.g., Queen, U.S. Patent No. 5,585,089, which is incorporated herein by reference in its entirety.) An immunoglobulin light or heavy chain variable region consists of a "framework" region interrupted by three hypervariable regions, referred to as complementarily determining regions (CDRs). The extent of the framework region and CDRs have been precisely defined (See, "Sequences of Proteins of Immunological Interest", Kabat, E. et al., U.S. Department of Health and Human Services (1983)). Briefly, humanized antibodies are antibody molecules from non-human species having one or more CDRs from the non-human species and a framework from a human immunoglobulin molecule.

The antibodies of the invention may also be labelled in the same manner as described above for the nucleic probes of the invention such as an enzymatic, fluorescent or radioactive type labelling.

The invention relates, in addition, to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:

- a) bringing the biological sample (biological tissue or fluid) into contact with a mono- or polyclonal antibody according to the invention (under conditions allowing an immunological reaction between the said antibodies and the polypeptides of the bacterium belonging to the species *Chlamydia trachomatis* or to an associated microorganism which may be present in the biological sample, that is, under conditions suitable for the formation of immune complexes):
 - b) detecting the antigen-antibody complex which may be formed.

Also falling within the scope of the invention is a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a polyclonal or monoclonal antibody according to the invention, labelled where appropriate;
- where appropriate, a reagent for constituting the medium appropriate for carrying out the immunological reaction;
 - a reagent allowing the detection of the antigen-antibody complexes produced by the immunological reaction, it being possible for this reagent also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the said monoclonal or polyclonal antibody is not labelled;
- 35 where appropriate, reagents for carrying out the lysis of the cells in the sample tested.

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The principle of the DNA chip which was explained above may also be used to produce protein «chips» on which the support has been coated with a polypeptide or an antibody according to the invention, or arrays thereof, in place of the DNA. These protein «chips» make it possible, for example, to analyse the biomolecular interactions (BIA) induced by the affinity capture of target analytes onto a support coated, for example, with proteins, by surface plasma resonance (SPR). Reference may be made, for example, to the techniques for coupling proteins onto a solid support which are described in EP 524 800 or to the methods describing the use of biosensor-type protein chips such as the BIAcore-type technique (Pharmacia) (Arlinghaus et al., 1997, Krone et al., 1997, Chatelier et al., 1995). These polypeptides or antibodies according to the invention, capable of specifically binding antibodies or polypeptides derived from the sample to be analysed, may thus be used in protein chips for the detection and/or the identification of proteins in samples. The said protein chips may in particular be used for infectious diagnosis and may preferably contain, per chip, several polypeptides and/or antibodies of the invention of different specificity, and/or polypeptides and/or antibodies capable of recognizing microorganisms different from Chlamydia trachomatis.

Accordingly, the subject of the present invention is also the polypeptides and the antibodies according to the invention, characterized in that they are immobilized on a support, in particular of a protein chip.

The protein chips, characterized in that they contain at least one polypeptide or one antibody according to the invention immobilized on the support of the said chip, also form part of the 20 invention.

The invention comprises, in addition, a protein chip according to the invention, characterized in that it contains, in addition, at least one polypeptide of a microorganism different from *Chlamydia trachomatis* or at least one antibody directed against a compound of a microorganism different from *Chlamydia trachomatis*, immobilized on the support of the said chip.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a protein chip according to the invention.

The subject of the present invention is also a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it uses a nucleotide sequence according to the invention.

More particularly, the invention relates to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:

- a) where appropriate, isolation of the DNA from the biological sample to be analysed, or optionally production of a cDNA from the RNA in the biological sample;
- b) specific amplification of the DNA of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism with the aid of at least one primer according to the invention;
- 5 c) detection of the amplification products.

These may be detected, for example, by the molecular hybridization technique using a nucleic probe according to the invention. This probe will be advantageously labelled with a nonradioactive (cold probe) or radioactive element.

For the purposes of the present invention, «DNA in the biological sample» or «DNA contained in the biological sample» will be understood to mean either the DNA present in the biological sample considered, or optionally the cDNA obtained after the action of a reverse transcriptase-type enzyme on the RNA present in the said biological sample.

Another aim of the present invention consists in a method according to the invention, characterized in that it comprises the following steps:

- a) bringing a nucleotide probe according to the invention into contact with a biological sample, the DNA contained in the biological sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to complementary base pairs of the DNA of a bacterium belonging to the species Chlamydia trachomatis or to an associated microorganism;
- 20 b) detecting the hybridization complex formed between the nucleotide probe and the DNA in the biological sample.

The present invention also relates to a method according to the invention, characterized in that it comprises the following steps:

- a) bringing a nucleotide probe immobilized on a support according to the invention into contact
 with a biological sample, the DNA in the sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to the DNA of a bacterium belonging to the species *Chlamydia trachomatis* or to an associated microorganism;
- b) bringing the hybrid formed between the nucleotide probe immobilized on a support and the DNA contained in the biological sample, where appropriate after removal of the DNA in the
 30 biological sample which has not hybridized with the probe, into contact with a labelled nucleotide probe according to the invention;
 - c) detecting the new hybrid formed in step b).

According to an advantageous embodiment of the method for the detection and/or the identification defined above, it is characterized in that, prior to step a), the DNA in the biological sample is primer-extended and/or amplified beforehand with the aid of at least one primer according

to the invention.

The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- 5 a) a nucleotide probe according to the invention;
 - b) where appropriate, the reagents necessary for carrying out a hybridization reaction;
 - c) where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a) a nucleotide probe, called capture probe, according to the invention;
- b) an oligonucleotide probe, called detection probe, according to the invention;
- c) where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a) at least one primer according to the invention:
- 20 b) where appropriate, the reagents necessary for carrying out a DNA amplification reaction;
 - c) where appropriate, a component which makes it possible to check the sequence of the amplified fragment, more particularly an oligonucleotide probe according to the invention.

The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a DNA chip according to the invention.

The invention also relates to a method or to a kit or set according to the invention for the detection and/or the identification of bacteria belonging to the species Chlamydia trachomatis, characterized in that the said primer and/or the said probe according to the invention are chosen from the nucleotide sequences specific to the species Chlamydia trachomatis, in that the said polypeptides according to the invention are chosen from the polypeptides specific to the species Chlamydia trachomatis and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides specific to the species Chlamydia trachomatis.

Preferably, the said method or the said kit or set above according to the invention, for

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the detection and/or the identification of bacteria belonging to the species Chlamydia trachomatis is characterized in that the said primer and/or the said probe or the said polypeptides are chosen from the nucleotide sequences or polypeptides according to the invention which have been identified as being specific to the species Chlamydia trachomatis and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides identified as being specific to the species Chlamydia trachomatis.

The invention relates, in addition, to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of a condition caused by, genital diseases which are induced or worsened by a *Chlamydia trachomatis* infection.

The invention also relates to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of conditions caused by, eye diseases induced or worsened by a *Chlamydia trachomatis* infection.

The invention also relates to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of conditions caused by, systemic diseases, in particular of the lymphatic system, which are induced or worsened by a *Chlamydia trachomatis* infection.

According to another aspect, the subject of the invention is the use of polypeptides according to the invention, of cells transformed with a vector according to the invention and/or of transformed animals according to the invention, for the biosynthesis or the biodegradation of organic or inorganic compounds.

As has been mentioned above, the nucleotide sequences of the invention were identified by homology with sequences known to encode, for example, polypeptides or fragments of enzymatic polypeptides involved in the biosynthesis or the biodegradation of organic or inorganic molecules.

It is thus possible to use the said polypeptides of the invention in a similar manner for the biosynthesis or the biodegradation of organic or inorganic compounds of industrial or therapeutic interest (called compounds of interest).

Among these polypeptides, there may be mentioned in particular the enzymes involved in metabolism, such as the proteolytic enzymes, amino transferases, glucose metabolism, or the enzymes which may be used in the biosynthesis of sugars, amino acids, fatty acids, polypeptides, nucleotides, nucleic acids or any other organic or inorganic compound or in the biodegradation of organic or inorganic compounds.

Among these polypeptides, there may be mentioned, in addition, the mutated or modified enzymes corresponding to mutated or modified polypeptides according to the invention which may also be used for the biosynthesis or the biodegradation of organic or inorganic compounds at the industrial level, such as, for example, the production of compounds of interest, the reprocessing

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of manufacturing residues applied to the food industries, to the papermaking industry or to the chemical and pharmaceutical industries.

The methods of biosynthesis or biodegradation of organic or inorganic compounds, characterized in that they use a polypeptide or one of its representative fragments according to the invention, transformed cells according to the invention and/or a transformed animal according to the invention, also form part of the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, of a polypeptide according to the invention, of an antibody according to the invention, of a cell according to the invention, and/or of a transformed animal according to the invention, for the selection of an organic or inorganic compound capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the cellular replication of eukaryotic or prokaryotic cells or capable of inducing, inhibiting or worsening the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms.

The invention also comprises screening assays that comprise method of selecting compounds capable of binding to a polypeptide, fusion polypeptide, or one of its representative fragments according to the invention, capable of binding to a nucleotide sequence according to the invention, or capable of recognizing an antibody according to the invention, and/or capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the growth or the cellular replication of eukaryotic or prokaryotic cells, or capable of inducing, inhibiting or worsening, in an animal or human organism, the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms, characterized in that it comprises the following steps:

- a) bringing the said compound into contact with the said polypeptide, the said nucleotide sequence, with a transformed cell according to the invention and/or administering the said compound to a transformed animal according to the invention;
- b) determining the capacity of the said compound to bind with the said polypeptide or the said nucleotide sequence, or to modulate, regulate, induce or inhibit the expression of genes, or to modulate growth or cellular replication, or to induce, inhibit or worsen in the said transformed animal, the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms.

The transformed cells and/or animals according to the invention may advantageously serve as a model and may be used in methods for studying, identifying and/or selecting compounds capable of being responsible for pathologies induced or worsened by *Chlamydia trachomatis*, or capable of preventing and/or of treating these pathologies such as, for example, genital, eye or systemic diseases, especially of the lymphatic system. In particular, the transformed host cells, in

particular bacteria of the *Chlamydia* family whose transformation with a vector according to the invention may, for example, increase or inhibit its infectivity, or modulate the pathologies usually induced or worsened by the infection, may be used to infect animals in which the onset of pathologies will be monitored. These nontransformed animals, infected for example with transformed *Chlamydia* bacteria, may serve as a study model. In the same manner, the transformed animals according to the invention may, for example, exhibit predispositions to genital and/or eye and/or systemic diseases, especially of the lymphatic system, and thus be used in methods for selecting compounds capable of preventing and/or of treating the said diseases. The said methods using the said transformed cells and/or transformed animals form part of the invention.

The compounds capable of being selected may be organic compounds such as polypeptides or carbohydrates or any other organic or inorganic compounds already known, or new organic compounds produced using molecular modelling techniques and obtained by chemical or biochemical synthesis, these techniques being known to persons skilled in the art.

The said selected compounds may be used to modulate the growth and/or the cellular replication of *Chlamydia trachomatis* or any other associated microorganism and thus to control infection by these microorganisms. The said compounds according to the invention may also be used to modulate the growth and/or the cellular replication of all eukaryotic or prokaryotic cells, in particular tumour cells and infectious microorganisms, for which the said compounds will prove active, the methods which make it possible to determine the said modulations being well known to persons skilled in the art.

Compound capable of modulating the growth of a microorganism is understood to designate any compound which makes it possible to act, to modify, to limit and/or to reduce the development, the growth, the rate of proliferation and/or the viability of the said microorganism.

This modulation may be achieved, for example, by an agent capable of binding to a protein and thus of inhibiting or of potentiating its biological activity, or capable of binding to a membrane protein of the outer surface of a microorganism and of blocking the penetration of the said microorganism into the host cell or of promoting the action of the immune system of the infected organism directed against the said microorganism. This modulation may also be achieved by an agent capable of binding to a nucleotide sequence of a DNA or RNA of a microorganism and of blocking, for example, the expression of a polypeptide whose biological or structural activity is necessary for the growth or for the reproduction of the said microorganism.

Associated microorganism is understood to designate in the present invention any microorganism whose gene expression may be modulated, regulated, induced or inhibited, or whose growth or cellular replication may also be modulated by a compound of the invention. Associated microorganism is also understood to designate in the present invention any microorganism containing

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nucleotide sequences or polypeptides according to the invention. These microorganisms may, in some cases, contain polypeptides or nucleotide sequences identical or homologous to those of the invention may also be detected and/or identified by the detection and/or identification methods or kit according to the invention and may also serve as a target for the compounds of the invention.

The invention relates to the compounds capable of being selected by a method of selection according to the invention.

The invention also relates to a pharmaceutical composition comprising a compound chosen from the following compounds:

a nucleotide sequence according to the invention;

10 a polypeptide or fusion polypeptide according to the invention;

a vector according to the invention;

an antibody according to the invention; and

a compound capable of being selected by a method of selection according to the invention, optionally in combination with a pharmaceutically acceptable vehicle or carrier.

An effective quantity is understood to designate a sufficient quantity of the said compound or antibody, or of a polypeptide of the invention, which makes it possible to modulate the growth of *Chlamydia trachomatis* or of an associated microorganism.

The invention also relates to a pharmaceutical composition according to the invention for the prevention or the treatment of an infection by a bacterium belonging to the species *Chlamydia trachomatis* or by an associated microorganism.

The invention relates, in addition, to an immunogenic and/or vaccine composition, characterized in that it comprises one or more polypeptides according to the invention and/or one or more hybrid polypeptides according to the invention.

The invention also comprises the use of a transformed cell according to the invention, for the preparation of a vaccine composition.

The invention also relates to a vaccine composition, characterized in that it contains a nucleotide sequence according to the invention, a vector according to the invention and/or a transformed cell according to the invention.

The invention also relates to the vaccine compositions according to the invention, for the prevention or the treatment of an infection by a bacterium belonging to the species *Chlamydia trachomatis* or by an associated microorganism.

The invention also relates to the use of DNA encoding polypeptides of *Chlamydia trachomatis*, in particular antigenic determinants, to be formulated as vaccine compositions. In accordance with this aspect of the invention, the DNA of interest is engineered into an expression vector under the control of regulatory elements, which will promote expression of the DNA, *i.e.*,

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promoter or enhancer elements. In one preferred embodiment, the promoter element may be cellspecific and permit substantial transcription of the DNA only in predetermined cells. The DNA may be introduced directly into the host either as naked DNA (U.S. Patent No. 5,679,647 incorporated herein by reference in their entirety) or formulated in compositions with other agents which may facilitate uptake of the DNA including viral vectors, i.e., adenovirus vectors, or agents which facilitate immunization, such as bupivicaine and other local anesthetics (U.S. Patent 5,593,972 incorporated herein by reference in their entirety), saponins (U.S. Patent 5,739,118 incorporated herein by reference in their entirety) and cationic polyamines (published international application WO 96/10038 incorporated herein by reference in their entirety).

The DNA sequence encoding the antigenic polypeptide and regulatory element may be inserted into a stable cell line or cloned microorganism, using techniques, such as targeted homologous recombination, which are well known to those of skill in the art, and described e.g., in Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

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Such cell lines and microorganisms may be formulated for vaccine purposes. In yet another embodiment, the DNA sequence encoding the antigenic polypeptide and regulatory element may be delivered to a mammalian host and introduced into the host genome via homologous recombination (See, Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

Preferably, the immunogenic and/or vaccine compositions according to the invention intended for the prevention and/or the treatment of an infection by Chlamydia trachomatis or by an associated microorganism will be chosen from the immunogenic and/or vaccine compositions comprising a polypeptide or one of its representative fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia trachomatis. The vaccine 25 compositions comprising nucleotide sequences will also preferably comprise nucleotide sequences encoding a polypeptide or one of its fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia trachomatis.

Among these preferred immunogenic and/or vaccine compositions, the most preferred are those comprising a polypeptide or one of its representative fragments, or a nucleotide sequence or one of its representative fragments whose sequences are chosen from the nucleotide or amino acid sequences identified in this functional group and listed above.

The polypeptides of the invention or their representative fragments entering into the immunogenic compositions according to the invention may be selected by techniques known to persons skilled in the art, such as for example on the capacity of the said polypeptides to stimulate T 35 cells, which results, for example, in their proliferation or the secretion of interleukins, and which leads to the production of antibodies directed against the said polypeptides.

In mice, in which a weight dose of the vaccine composition comparable to the dose used in humans is administered, the antibody reaction is tested by collecting serum followed by a study of the formation of a complex between the antibodies present in the serum and the antigen of the vaccine composition, according to the customary techniques.

According to the invention, the said vaccine compositions will be preferably in combination with a pharmaceutically acceptable vehicle and, where appropriate, with one or more appropriate immunity adjuvants.

Various types of vaccines are currently available for protecting humans against infectious diseases: attenuated live microorganisms (M. bovis - BCG for tuberculosis), inactivated microorganisms (influenza virus), acellular extracts (Bordetella pertussis for whooping cough), recombinant proteins (hepatitis B virus surface antigen), polysaccharides (pneumococci). Experiments are underway on vaccines prepared from synthetic peptides or from genetically modified microorganisms expressing heterologous antigens. Even more recently, recombinant plasmid DNAs carrying genes encoding protective antigens were proposed as an alternative vaccine strategy. This type of vaccination is carried out with a particular plasmid derived from an E. coli plasmid which does not replicate in vivo and which encodes only the vaccinal protein. Animals were immunized by simply injecting the naked plasmid DNA into the muscle. This technique leads to the expression of the vaccine protein in situ and to a cell-type (CTL) and a humoral type (antibody) immune response.

This double induction of the immune response is one of the main advantages of the technique of vaccination with naked DNA.

The vaccine compositions of the present invention can be evaluated in *in vitro* and *in vivo* animal models prior to host, e.g., human, administration. For example, *in vitro* neutralization assays such as those described by Peterson et al. (1988) can be utilized. The assay described by Peterson et al. (1988) is suitable for testing vaccine compositions directed toward either *Chlamydia trachomatis* or *Chlamydia pneumoniae*.

Briefly, hyper-immune antisera is diluted in PBS containing 5% guinea pig serum, as a complement source. Chlamydiae (10⁴ IFU; inclusion forming units) are added to the antisera dilutions. The antigen-antibody mixtures are incubated at 37°C for 45 minutes and inoculated into duplicate confluent Hep-2 or HeLa cell monolayers contained in glass vials (e.g., 15 by 45 mm), which have been washed twice with PBS prior to inoculation. The monolayer cells are infected by centrifugation at 1000X g for 1 hour followed by stationary incubation at 37° for 1 hour. Infected monolayers are incubated for 48 or 72 hours, fixed and stained with a Chlamydiae specific antibody, such as anti-MOMP for C. trachomatis, etc. Inclusion-bearing cells are counted in ten fields at a magnification of 200X. Neutralization titer is assigned based on the dilution that gives 50%

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inhibition as compared to control monolayers/IFU.

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The efficacy of vaccine compositions can be determined *in vivo* by challenging animal models of *Chlamydia trachomatis* infection, *e.g.*, guinea pigs or mice, with the vaccine compositions. For example, *in vivo* vaccine composition challenge studies in the guinea pig model of *Chlamydia trachomatis* infection can be performed. Briefly, female guinea pigs weighing 450 to 500 g are housed in an environmentally controlled room with a 12 hour light-dark cycle and immunized with vaccine compositions via a variety of immunization routes. Post-vaccination, guinea pigs are infected in the genital tract with the agent of guinea pig inclusion conjunctivitis (GPIC), which has been grown in HeLa or McCoy cells (Rank et al. (1988)). Each animal receives approximately 1.4x10⁷ inclusion forming units (IFU) contained in 0.05 ml of sucrose-phosphate-glutamate buffer, pH 7.4 (Schacter, J. (1980)). The course of infection monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorescence with GPIC specific antisera, or by Giernsastained smear from a scraping from the genital tract (Rank et al. (1988)). Antibody titers in the serum is determined by an enzyme-linked immunosorbent assay.

Alternatively, in vivo vaccine composition challenge studies can be performed in the murine model of Chlamydia trachomatis (Morrison et al., 1995). Briefly, female mice 7 to 12 weeks of age receive 2.5 mg of depoprovera subcutaneously at 10 and 3 days before vaginal infection. Post-vaccination, mice are infected in the genital tract with 1,500 inclusion-forming units of Chlamydia trachomatis contained in 5ml of sucrose-phosphate-glutamate buffer, pH. 7.4. The course of infection is monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorence with Chlamydia trachomatis specific antisera, or by a Giemsa-stained smear from a scraping from the genital tract of an infected mouse. The presence of antibody titers in the serum of a mouse is determined by an enzyme-linked immunosorbent assay.

The vaccine compositions comprising nucleotide sequences or vectors into which the said sequences are inserted are in particular described in International Application No. WO 90/11092 and also in International Application No. WO 95/11307.

The nucleotide sequence constituting the vaccine composition according to the invention may be injected into the host after having been coupled to compounds which promote the penetration of this polynucleotide inside the cell or its transport up to the cell nucleus. The resulting conjugates may be encapsulated into polymeric microparticles, as described in International Application No. WO 94/27238 (Medisorb Technologies International).

According to another embodiment of the vaccine composition according to the invention, the nucleotide sequence, preferably a DNA, is complexed with the DEAE-dextran (Pagano et al., 1967) or with nuclear proteins (Kaneda et al., 1989), with lipids (Felgner et al., 1987) or encapsulated into liposomes (Fraley et al., 1980) or alternatively introduced in the form of a gel

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facilitating its transfection into the cells (Midoux et al., 1993, Pastore et al., 1994). The polynucleotide or the vector according to the invention may also be in suspension in a buffer solution or may be combined with liposomes.

Advantageously, such a vaccine will be prepared in accordance with the technique described by Tacson et al. or Huygen et al. in 1996 or alternatively in accordance with the technique described by Davis et al. in International Application No. WO 95/11307.

Such a vaccine may also be prepared in the form of a composition containing a vector according to the invention, placed under the control of regulatory elements allowing its expression in humans or animals. It is possible, for example, to use, as vector for the *in vivo* expression of the polypeptide antigen of interest, the plasmid pcDNA3 or the plasmid pcDNA1/neo, both marketed by Invitrogen (R & D Systems, Abingdon, United Kingdom). It is also possible to use the plasmid V1Jns.tPA, described by Shiver et al. in 1995. Such a vaccine will advantageously comprise, in addition to the recombinant vector, a saline solution, for example a sodium chloride solution.

The immunogenic compositions of the invention can be utilized as part of methods of immunization, wherein such methods comprise administering to a host, e.g., a human host, an immunizing amount of the immunogenic compositions of the invention. In a preferred embodiment, the method of immunizing is a method of immunizing against *Chlamydia trachomatis*.

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A pharmaceutically acceptable vehicle is understood to designate a compound or a combination of compounds entering into a pharmaceutical or vaccine composition which does not cause side effects and which makes it possible, for example, to facilitate the administration of the active compound, to increase its life and/or its efficacy in the body, to increase its solubility in solution or alternatively to enhance its preservation. These pharmaceutically acceptable vehicles are well known and will be adapted by persons skilled in the art according to the nature and the mode of administration of the active compound chosen.

As regards the vaccine formulations, these may comprise appropriate immunity adjuvants which are known to persons skilled in the art, such as, for example, aluminium hydroxide, a representative of the family of muramyl peptides such as one of the peptide derivatives of N-acetyl-muramyl, a bacterial lysate, or alternatively incomplete Freund's adjuvant, StimulonTM QS-21 (Aquila Biopharmaceuticals, Inc., Framinham, MA), MPLTM (3-0-deacylated monophosphoryl lipid A; RIBI ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA).

Preferably, these compounds will be administered by the systemic route, in particular by the intravenous route, by the intranasal, intramuscular, intradermal or subcutaneous route, or by the oral route. More preferably, the vaccine composition comprising polypeptides according to the invention will be administered several times, spread out over time, by the intradermal or subcutaneous

route.

Their optimum modes of administration, dosages and galenic forms may be determined according to criteria which are generally taken into account in establishing a treatment adapted to a patient, such as for example the patient's age or body weight, the seriousness of his general condition, tolerance of the treatment and the side effects observed.

The invention comprises the use of a composition according to the invention for the treatment or the prevention of genital diseases which are induced or worsened by *Chlamydia trachomatis*.

Finally, the invention comprises the use of a composition according to the invention for the treatment or the prevention of eye diseases which are induced or worsened by the presence of *Chlamydia trachomatis*.

Finally, the invention comprises the use of a composition according to the invention for the treatment or the prevention of systemic diseases, especially of the lymphatic system, which are induced or worsened by the presence of *Chlamydia trachomatis*.

Other characteristics and advantages of the invention appear in the following examples and figures:

Legend to the figures:

Figure 1: Line for the production of Chlamydia trachomatis sequences

20 Figure 2: Analysis of the sequences and assembling

Figure 3: Finishing techniques

Figure 3a): Assembly map

Figure 3b): Determination and use of the orphan ends of the contigs

25 **EXAMPLES**

Cells

The Chlamydia trachomatis LGV2 strain used is identified to have over 98% homology with the outer membrane protein sequences omp1 (CHTMOMPA) and omp2 (CHTOMP2A) of the Chlamydia trachomatis serovar L2/434/Bu strain.

The Chlamydia trachomatis LGV2 strain is cultured on mouse fibroblasts (McCoy cells), obtained from the American Type Culture Collection, under the reference ATCC CRL-1696.

Culture of the cells

The mouse fibroblasts are cultured in 75-ml cell culture flasks (Corning). The culture medium is Dulbecco's modified cell culture medium (Gibco BRL No. 04101965) supplemented with

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MEM amino acids (Gibco BRL - No. 04301140) L (5 ml per 500 ml of medium) and 5% foetal calf serum (Gibco BRL No. 10270 batch 40G8260K) without antibiotics or antifungals.

The cell culture stock is maintained in the following manner. The cell cultures are examined under an inverted microscope. 24 hours after confluence, each cellular lawn is washed with PBS (Gibco BRL No. 04114190), rinsed and then placed for 5 min in an oven in the presence of 3 ml of trypsine (Gibco BRL No. 25200056). The cellular lawn is then detached and then resuspended in 120 ml of culture medium, the whole is stirred in order to make the cellular suspension homogeneous. 30 ml of this suspension are then distributed per cell culture flask. The flasks are kept in a CO₂ oven (5%) for 48 hours at a temperature of 37°C. The cell stock is maintained so as to have available daily 16 flasks of subconfluent cells. It is these subconfluent cells which will be used so as to be infected with Chlamydia. 25-ml cell culture flasks are also used, these flasks are prepared in a similar manner but the volumes used for maintaining the cells are the following: 1 ml of trypsine, 28 ml of culture medium to resuspend the cells, 7 ml of culture medium are used per 25-ml flask.

15 <u>Infection of the cells with Chlamydia</u>

Initially, the Chlamydiae are obtained frozen (at -70°C), in suspension in a volume of 1 millilitre. This preparation is slowly thawed, 500 µl are collected and brought into contact with subconfluent cells, which are obtained as indicated above, in a 25-ml cell culture flask, containing 1 ml of medium, so as to cover the cells. The flask is then centrifuged at 2000 rpm in a «swing» rotor for microtitre plates, the centrifuge being maintained at a temperature of 35°C. After centrifugation, the two flasks are placed in an oven at 35°C for three hours. 6 ml of culture medium containing cycloheximide (1 µg/ml) are then added and the flask is stored at 35°C. After 48 hours, the level of infection is evaluated by direct immunofluorescence and by the cytopathogenic effect caused to the cells.

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Direct immunofluorescence

Starting with infected cells, which were obtained as indicated above, a cellular smear is deposited with a Pasteur pipette on a microscope slide. The cellular smear is fixed with acetone for 10 minutes; after draining the acetone, the smear is covered with 30 µl of murine monoclonal antibodies directed against MOMP (major outer membrane protein) of Chlamydia (Syva, Biomérieux) labelled with fluorescein isothiocyanate. The whole is then incubated in a humid chamber at a temperature of 37°C. The slides are then rinsed with water, slightly dried, and then after depositing a drop of mounting medium, a coverslip is mounted before reading. The reading is carried out with the aid of a fluorescence microscope equipped with the required filters (excitation at 490 nm, emission at 520 nm).

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Harvesting of the Chlamydia trachomatis

After checking the infection by direct immunofluorescence, carried out as indicated above, the culture flasks are opened under a sterile cabinet, sterile glass beads with a diameter of the 5 order of a millimeter are placed in the flask. The flask is closed and then vigorously stirred while being maintained horizontally, the cellular lawn at the bottom, so that the glass beads can have a mechanical action on the cellular lawn. Most of the cells are thus detached or broken; the effect of the stirring is observed under an optical microscope so as to ensure proper release of Chlamydiae.

Large-scale infection of the cell cultures

The product of the Chlamydiae harvest (culture medium and cellular debris) is collected with a pipette, and distributed into three cell culture flasks containing subconfluent L cells, obtained as indicated above. The cells thus inoculated are placed under gentle stirring (swing) in an oven at 35°C. After one hour, the flasks are kept horizontally in an oven so that the culture medium covers the cells for 3 hours. 30 ml of culture medium containing actydione (1 µg/ml) are then added to each of the flasks. The culture flasks are then stored at 35°C for 48 hours. The cells thus infected are examined under an optical microscope after 24 hours, the cytopathogenic effect is evaluated by the appearance of cytoplasmic inclusions which are visible under an inverted optical microscope. After 48 hours, the vacuoles containing the Chlamydiae occupy the cytoplasm of the cell and push the cell nucleus sideways. At this stage, numerous cells are spontaneously destroyed and have left free elementary bodies in the culture medium. The Chlamydiae are harvested as described above and are either frozen at -80°C or used for another propagation.

Purification of the Chlamydiae

The product of the Chlamydia harvests, stored at -80°C, is thawed on a water bath at room temperature. After thawing, each tube is vigorously stirred for one minute and immersed for one minute in an ultrasound tank (BRANSON 1200); the tubes are then stirred by inverting before being centrifuged for 5 min at 2000 rpm. The supernatant is carefully removed and kept at cold temperature (ice). The supernatant is vigorously stirred and then filtered on nylon filters having pores of 5 microns 30 in diameter on a support (Nalgene) allowing a delicate vacuum to be established under the nylon filter. For each filtration, three nylon filters are superposed; these filters are replaced after every 40 ml of filtrate. Two hundred milliliters of filtration product are kept at cold temperature, and then after stirring by inverting, are centrifuged at 10,000 rpm for 90 min, the supernatant is removed and the pellet is taken up in 10 ml of 10 mM Tris, vigorously vortexed and then centrifuged at 10,000 rpm for 35 90 min. The supernatant is removed and the pellet is taken up in a buffer (20 mM Tris pH 8.0, 50 mM

KCl, 5 mM MgCl₂) to which 800 units of DNAse I (Boehringer) are added. The whole is kept at 37°C for one hour. One ml of 0.5 M EDTA is then added, and the whole is vortexed and frozen at -20°C.

Preparation of the DNA

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The Chlamydiae purified above are thawed and subjected to a proteinase K (Boehringer) digestion in a final volume of 10 ml. The digestion conditions are the following: 0.1 mg/ml proteinase K, 0.1 % SDS at 55°C, stirring every 10 min. The product of digestion is then subjected to a double extraction with phenol-chloroform, two volumes of ethanol are added and the DNA is directly recovered with a Pasteur pipette having one end in the form of a hook. The DNA is 10 dried on the edge of the tube and then resuspended in 500 µl of 2 mM Tris pH 7.5. The DNA is stored at 4°C for at least 24 hours before being used for the cloning.

Cloning of the DNA

After precipitation, the DNA is quantified by measuring the optical density at 260 nm. Thirty μg of Chlamydia DNA are distributed into 10 tubes of 1.5 ml and diluted in 300 μl of water. Each of the tubes is subjected to 10 applications of ultrasound lasting for 0.5 sec in a sonicator (Unisonix XL2020). The contents of the 10 tubes are then grouped and concentrated by successive extractions with butanol (Sigma B1888) in the following manner: two volumes of butanol are added to the dilute DNA mixture. After stirring, the whole is centrifuged for five minutes at 2500 rpm and the butanol is removed. This operation is repeated until the volume of the aqueous phase is less than 1 ml. The DNA is then precipitated in the presence of ethanol and of 0.5 M sodium acetate pH 5.4, and then centrifuged for thirty minutes at 15,000 rpm at cold temperature (4°C). The pellet is washed with 75% ethanol, centrifuged for five minutes at 15,000 rpm and dried at room temperature. A tenth of the preparation is analysed on a 0.8% agarose gel. Typically, the size of the DNA fragments thus prepared is between 200 and 8000 base pairs.

To allow the cloning of the DNA obtained, the ends are repaired. The DNA is distributed in an amount of 10 µg/tube, in the following reaction medium: 100 µl final volume, 1 H buffer (Biolabs 201L), 0.5 µl BSA 0.05 mg/ml, 0.1 mM dATP, 0.1 mM each of dGTP, dCTP or dTTP, 60,000 IU T4 DNA polymerase. The reaction is incubated for thirty minutes at 16°C. The contents of each of the tubes are then grouped before carrying out an extraction with phenolchloroform and then precipitating the aqueous phase as described above. After this step, the DNA thus prepared is phosphorylated. For that, the DNA is distributed into tubes in an amount of 10 µg per tube, and then in a final volume of 50 µl, the reaction is prepared in the following manner: 1 mM ATP, 1 x kinase buffer, 10 IU T4 polynucleotide kinase (Biolabs 201L). The preparation is incubated for thirty minutes at 37°C. The contents of the tubes are combined and a phenol-chloroform extraction

and then a precipitation are carried out in order to precipitate the DNA. The latter is then suspended in 1 µl of water and then the DNA fragments are separated according to their size on a 0.8% agarose gel (1 x TAE). The DNA is subjected to an electric field of 5 V/cm and then visualized on a UV table. The fragments whose size varies between 1200 and 2000 base pairs are selected by cutting out the gel.

The gel fragment thus isolated is placed in a tube and then the DNA is purified with the Qiaex kit (20021 Qiagen), according to the procedure provided by the manufacturer.

Preparation of the vector

14 μg of the cloning vector pGEM-5Zf (Proméga P2241) are diluted in a final volume of 150 μl and are subjected to digestion with the restriction enzyme EcoRV 300 IU (Biolabs 195S) according to the protocol and with the reagents provided by the manufacturer. The whole is placed at 37°C for 150 min and then distributed in the wells of a 0.8% agarose gel subjected to an electric field of 5 V/cm. The linearized vector is visualized on a UV table, isolated by cutting out the gel and then purified by the Qiaex kit (Qiagen 20021) according to the manufacturer's recommendations. The purification products are grouped in a tube, the volume is measured and then half the volume of phenol is added and the whole is vigorously stirred for 1 min. Half the volume of chloroform-isoamyl alcohol 24:1 is added and vigorously stirred for 1 min. The whole is centrifuged at 15,000 rpm for 5 min at 4°C, the aqueous phase is recovered and transferred into a tube. The DNA is precipitated in the presence of 0.3 M sodium acetate, pH 5.4 and 3 volumes of ethanol and placed at -20°C for 1 hour. The DNA is then centrifuged at 15,000 rpm for 30 min at 4°C, the supernatant is removed while preserving the pellet, washed twice with 70% ethanol. After drying at room temperature, the DNA is suspended in 25 μl of water.

Phosphorylation of the vector

25 μl of the vector prepared in the preceding step are diluted in a final volume of 500 μl of the following reaction mixture:

After repair, the DNA is subjected to a phenol-chloroform extraction and a precipitation, the pellet is then taken up in 10 µl of water, the DNA is quantified by measuring the optical density at 260 nm. The quantified DNA is ligated into the vector PGEm-5Zf(+) prepared by the restriction enzyme EcoRV and dephosphorylated (see preparation of the vector). The ligation is carried out under three conditions which vary in the ratio between the number of vector molecules and the number of insert molecules. Typically, an equimolar ratio, a ratio of 1:3 and a ratio of 3:1 are used for the ligations which are, moreover, carried out under the following conditions: vector PGEm-5Zf(+) 25 ng, cut DNA, ligation buffer in a final volume of 20 µl with T4 DNA ligase (Amersham E70042X); the whole is then placed in a refrigerator overnight and then a phenol-

chloroform extraction and a precipitation are carried out in a conventional manner. The pellet is taken up in $5 \,\mu l$ of water.

Transformation of the bacteria

Plating of the bacteria

Petri dishes containing LB Agar medium containing ampicillin (50 µg/ml), Xgal (280 µg/ml) [5-bromo-4-chloro-indolyl-beta-D-galactopyranoside (Sigma B-4252)], IPTG (140 µg/ml) [isopropyl-beta-D-thiogalactoside (Sigma I-6758)] are used, 50 and 100 µl of bacteria are plated for each of the ligations. The Petri dishes are placed upside down at 37°C for 15 to 16 hours in an oven.

The number of «recombinant» positive clones is evaluated by counting the white colonies and the blue colonies which are thought to contain the vector alone.

Evaluation of the «recombinant» positive clones:

Ninety-four white colonies and two blue colonies are collected with the aid of sterile cones and are deposited at the bottom of the wells of plates designed for carrying out the amplification techniques. 30 µl of the following reaction mixture are added to each well: 1.7 mM MgCl₂, 0.2 mM each of dATP, dCTP, dGTP and dTTP, two synthetic oligonucleotides corresponding to sequences flanking the cloning site on either side and orienting the synthesis of the DNA in a convergent manner (0.5 µM RP and PU primers, 1 U TAQ polymerase (GibcoBRL 18038-026)).

The colonies thus prepared are subjected to a temperature of 94°C for 5 min and then to 30 thermal cycles composed of the following steps: 94°C for 40 s, 50°C for 30 s, 72°C for 180 s. The reaction is then kept for 7 min at 72°C and then kept at 4°C.

The amplification products are deposited on an agarose gel (0.8%), stained with ethidium bromide, subjected to electrophoresis, and then analysed on an ultraviolet table. The presence of an amplification fragment having a size greater than 500 base pairs indicates the presence of an insert. The bacterial clones are then prepared so as to study the sequence of their insert.

Sequencing

To sequence the inserts of the clones obtained as above, these were amplified by PCR on bacteria cultures carried out overnight using the primers for the vectors flanking the inserts. The sequence of the ends of these inserts (on average 500 bases on each side) was determined by automated fluorescent sequencing on an ABI 377 sequencer, equipped with the ABI Prism DNA Sequencing Analysis software (version 2.1.2).

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The sequences obtained by sequencing in a high-yield line (Figure 1) are stored in a database; this part of the production is independent of any treatment of the sequences. The sequences are extracted from the database, avoiding all the regions of inadequate quality, that is to say the regions for which uncertainties are observed on the sequence at more than 95%. After extraction, the 5 sequences are introduced into a processing line, the diagram of which is described in Figure 2. In a first path of this processing line, the sequences are assembled by the Gap4 software from R. Staden (Bonfield et al., 1995) (OS UNIX/SUN Solaris); the results obtained by this software are kept in the form of two files which will be used for a subsequent processing. The first of these files provides information on the sequence of each of the contigs obtained. The second file represents all the clones participating in the composition of all the contigs as well as their positions on the respective contigs.

The second processing path uses a sequence assembler (TIGR-Asmg assembler UNIX/SUN Solaris); the results of this second processing path are kept in the form of a file in the TIGR-Asmg format which provides information on the relationship existing between the sequences selected for the assembly. This assembler is sometimes incapable of linking contigs whose ends overlap over several hundreds of base pairs.

The results obtained from these two assemblers are compared with the aid of the BLAST program, each of the contigs derived from one assembly path being compared with the contigs derived from the other path.

For the two processing paths, the strict assembly parameters are fixed (95% homology, 30 superposition nucleotides). These parameters avoid 3 to 5% of the clones derived from eukaryotic cells being confused with sequences obtained from the clones derived from Chlamydia trachomatis. The eukaryotic sequences are however preserved during the course of this project; the strategy introduced, which is described below, will be designed, inter alia, not to be impeded by these sequences derived from contaminating clones.

The results of these two assemblers are processed in a software developed for this project. This software operates on a Windows NT platform and receives, as data, the results derived from the STADEN software and/or the results derived from the TIGR-Asmg assembler, the software, results, after processing of the data, in the determination of an assembly map which gives the proximity relationship and the orientation of the contigs in relation to one another (Figure 3a). Using 30 this assembly map, the software determines all the primers necessary for finishing the project. This treatment, which will be detailed below, has the advantage of distinguishing the isolated sequences derived from the contaminations, by the DNA eukaryotic cells, of the small-sized sequences clearly integrated into the project by the relationships which they establish with contigs. In order to allow, without any risk of error, the arrangement and the orientation of the contigs in relation to one another, a statistical evaluation of the accuracy of the names «naming» of sequence is made from the results of

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«contigation». This evaluation makes it possible to give each of the clone plates, as well as each of the subsets of plates, a weight which is inversely proportional to probable error rate existing in the «naming» of the sequences obtained from this plate or from a subset of this plate. In spite of a low error rate, errors may occur throughout the steps of production of the clones and of the sequences. These steps are numerous, repetitive and although most of them are automated, others, like the deposition in the sequencers, are manual; it is then possible for the operator to make mistakes such as the inversion of two sequences. This type of error has a repercussion on the subsequent processing of the data, by resulting in relationships (between the contigs) which do not exist in reality, then in attempts at directed sequencing between the contigs which will end in failure. It is because of this that 10 the evaluation of the naming errors is of particular importance since it allows the establishment of a probabilistic assembly map from which it becomes possible to determine all the clones which will serve as template to obtain sequences separating two adjacent contigs. Table 2 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997, each of which is 15 incorporated by reference herein in its entirety, gives the clones and the sequences of the primers initially used during the initial operations.

To avoid the step which consists in ordering and then preparing the clones by conventional microbiological means, outer and inner primers oriented towards the regions not yet sequenced are defined by the software. The primers thus determined make it possible to prepare, by 20 PCR, a template covering the nonsequenced region. It is the so-called outer primers (the ones most distant from the region to be sequenced) which are used to prepare this template. The template is then purified and a sequence is obtained on each of the two strands during 2 sequencing reactions which each use one of the 2 inner primers. In order to facilitate the use of this approach, the two outer primers and the two inner primers are prepared and then stored on the same position of 4 different 25 96-well plates. The two plates containing the outer primers are used to perform the PCRs which serve to prepare the templates. These templates will be purified on purification columns preserving the topography of the plates. Each of the sequences are obtained using primers situated on one and then on the other of the plates containing the inner primers. This distribution allows a very extensive automation of the process and results in a method which is simple to use for finishing the regions not 30 yet sequenced. Table 3 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997, each of which is incorporated by reference herein in its entirety, gives the names and the sequences of the primers used for finishing Chlamydia trachomatis.

Finally, a number of contigs exist in a configuration where one of their ends is not linked to any other contig end (Figure 3b) by a connecting clone relationship (a connecting clone is

defined as a clone having one sequence end on a contig and the other end of its sequence on another contig; furthermore, this clone must be derived from a plate or a subset of plates with adequate naming quality). For the Chlamydia trachomatis project, this particular case occurred 37 times. Two adjacent PCR primers orienting the synthesis of the DNA towards the end of the consensus sequence are defined for each of the orphan ends of the consensus sequence. The primer which is closest to the end of the sequence is called the inner primer whereas the primer which is more distant from the end of the sequence is called the outer primer. The outer primers are used to explore the mutual relationship between the orphan ends of the different contigs. The presence of a single PCR product and the possibility of amplifying this product unambiguously using the inner primers evokes the probable relationship between the contigs on which the primers which allowed the amplification are situated. This relationship will be confirmed by sequencing and will allow the connection between the orphan ends of the consensus sequences. This strategy has made it possible to obtain a complete map of the Chlamydia trachomatis chromosome and then to finish the project.

15 Quality control

All the bases not determined with certainty in the chromosomal sequence were noted and the density of uncertainties was measured on the entire chromosome. The regions with a high density of uncertainties were noted and the PCR primers spanning these regions were drawn and are represented in Table 4 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997 each of which is incorporated by reference herein in its entirety.

Data banks

Local reorganizations of major public banks were used. The protein bank used consists of the nonredundant fusion of the Genpept bank (automated translation of GenBank, NCBI; Benson et al., 1996).

The entire BLAST software (public domain, Altschul et al., 1990) for searching for homologies between a sequence and protein or nucleic data banks was used. The significance levels used depend on the length and the complexity of the region tested as well as the size of the reference bank. They were adjusted and adapted to each analysis.

The results of the search for homologies between a sequence according to the invention and protein or nucleic data banks are presented and summarized in Table 1 below.

Table 1: <u>List of coding chromosome regions and homologies between these regions and the sequence</u>

35 <u>banks.</u>

Legend to Table 1: Open reading frames are identified with the GenMark software version 2.3A (GenePro), the template used is *Chlamydia trachomatis* of order 4 on a length of 196 nucleotides with a window of 12 nucleotides and a minimum signal of 0.5. These reading frames are numbered in order of appearance on the chromosome, starting with ORF2 (ORF column). The positions of the beginning and of the end are then given in column 2 (position). When the position of the beginning is greater than the position of the end, this means that the region is encoded by the strand complementary to the sequence which was given in the sequence SEQ ID No. 1.

All the putative products were subjected to a search for homology on GENPEPT (release 103 for SEQ ID No. 2 to SEQ ID No. 1076 and release 108 for SEQ ID No. 1077 to SEQ ID No. 1197 with the BLASTp software (Altschul et al. 1990), with, as parameters, the default parameters with the exception of the expected value E set at 10⁻⁵ (for SEQ ID No. 2 to SEQ ID No. 1076) and P value set at e⁻¹⁰ (for SEQ ID No. 1077 to SEQ ID No. 1197). Subsequently, only the identities greater than 30% (I% column) were taken into account. The description of the most homologous sequence is given in the Homology column; the identifier for the latter sequence is given in the ID column and the animal species to which this sequence belongs is given in the Species column. The Homology score is evaluated by the sum of the blast scores for each region of homology and reported in the Score column. Table 1 also reflects data from additional ORF finder programs as defined below.

20 <u>Materials and methods: transmembrane domains:</u>

The DAS software was used as recommended by the authors (Cserzo et al., 1997).

This method uses, to predict the transmembrane domains, templates derived from a sampling of selected proteins. All the regions for which a "Cutoff" greater than 1.5 was found by the program were taken into account.

Additional ORF Finder Programs

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For this analysis, two additional ORF finder programs were used to predict potential open reading frames of a minimum length of 74 amino acids; Glimmer (Salzberg, S.L., Delcher, A., Kasif, S., and W. White. 1998. Microbial gene identification using interpolated Markov models.

Nucleic Acids Res. 26:544-548.), and an in-house written program. The in-house program used a very simple search algorithm. The analysis required the that the genomic DNA sequence text be in the 5' to 3' direction, the genome is circular, and that TAA, TAG, and TGA are stop codons. The search parameters were as follows:

(1) A search for an ORF that started with a GTG codon was performed. If no GTG codons were found, then a search for an ATG codon was performed. However, if a GTG codon was found,

then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.

- (2) A search for an ORF that started with a TTG codon was performed. If no TTG codons were found, then a search for a ATG codon was performed. However, if a TTG codon was found, 5 then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.
 - (3) The analysis described in steps 1 and 2 were repeated for the opposite strand of DNA sequence.
- (4) A search for ORFs that determined all ORF lengths using start and stop positions in the same reading frames was performed.
 - (5) All ORFs whose DNA length was less than 225 nucleotides were eliminated from the search.

Surface Exposed Protein Search Criteria

Potential cell surface vaccine targets are outer membrane proteins such as porins, lipoproteins, adhesions and other non-integral proteins. In Chlamydia psittaci, the major immunogens is a group of putative outer membrane proteins (POMPs) and no homologs have been found in Chlamydia trachomatis and Chlamydia trachomatis by traditional analysis (Longbottom, D., Russell, M., Dunbar, S.M., Jones, G.E., and A.J. Herring. 1998. Molecular Cloning and 20 Characterization of the Genes Coding for the Highly Immunogenic Cluster of 90-Kilodalton Envelope Proteins from Chlamydia psittaci Subtype That Causes Abortion in Sheep. Infect Immun 66:1317-1324.) However, utilizing the criteria described below, several ORFs encoding outer membrane proteins have been identified in Chlamydia trachomatis, all of which may represent vaccine candidates. Any ORF which met any one of the criteria described below were considered to encode a surface exposed protein.

Protein homology searches of the translated ORFs were done using the Blastp 2.0 tool (Altschul, S.F., Madden, T.L., Schaffer, A.A., Zhang, J., Zhang, Z., Miller, W., and D.J. Lipman. 1997. Gapped BLAST and PSI-BLAST: a new generation of protein database search programs. Nucleic Acids Res. 25:3389-3402). An ORF was labeled surface exposed if the translated ORF had 30 homology to a known, or hypothetical, or putative surface exposed protein with a P score less than e⁻¹⁰.

Most, if not all, proteins that are localized to the membrane of bacteria, via a secretory pathway, contain a signal peptide. The software program SignalP, was used to analyze the amino acid sequence of an ORF for such a signal peptide (Nielsen, H., Engelbrecht. J., Brunak, S., and G. von Heijne. 1997. Identification of prokaryotic and eukaryotic signal peptides and prediction

of their cleavage sites. Protein Engineering 10:1-6.) The first 60 N-terminal amino acids of each ORF were analyzed by SignalP using the Gram-negative software database. The output generates four separate values, maximum C, maximum Y, maximum S, and mean S. The S-score, or signal region, is the probability of the position belonging to the signal peptide. The C-score, or cleavage site, is the probability of the position being the first in the mature protein. The Y-score is the geometric average of the C-score and a smoothed derivative of the S-score. A conclusion of either a Yes or No is given next to each score. If all four conclusions are Yes and the C-terminal amino acid is either a phenylalanine (F) or a tyrosine (Y), the ORF was labelled outer membrane (Struyve, M., Moons, M., and J. Tommassen. 1991. Carboxy-terminal Phenylalanine is Essential for the Correct Assembly of a Bacterial Outer Membrane Protein. J. Mol. Biol. 218:141-148.)

The program called Psort was used to determine the localization of a protein based on its signal sequence, recognition of transmembrane segments, and analysis of its amino acid composition (Nakai, K., and M. Kanehisa. 1991. Expert system for predicting protein localization sites in gram-negative bacteria. Proteins 11:95-110.) An ORF is considered to be an outer membrane protein if the output data predicts the ORF encoded protein as outer membrane with a certainty value of 0.5 or better and whose value is at least twice as large as the next predicted localized certainty value.

Finally, ORFs that were not predicted to be outer membrane or surface exposed, based on the above criteria, were further analyzed. The Blastp output data for these ORFs were searched using various general and specific keywords, suggestive of known cell surface exposed proteins. An ORF was labeled surface exposed if the keywords matched had a Blastp hit with a P score less than e⁻¹⁰, and there was no better data indicating otherwise. The following is a list of the searched keywords:

25	Adhesion	Adhesin	Invasin
	Invasion	Extension	Omp
	Outer Surface	Porin	Outer Membrane
	Cell Surface	Cell Wall	Pilin
	Flagellar sheath	Cir	ChuA
30	СорВ	ExeD	FadL
	FecA	FepA	FhuA
	FmdC	FomA	FrpB
	GspD	HemR	HgbA
	Hgp	HmbR	HmuR
35	HMW	HrcC	Нгр

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	InvG	LamB	LbpA
	LcrQ	Lmp1	MxiD
	MOMP	PilE	HpaA
	NoIW	NspA	OpcP
5	OpnP	Opr	OspA
	PhoE	PldA	Por
	PscC	PulD	PupA
	QuiX	RafY	ScrY
	SepC	ShuA	SomA
10	SpiA	Tbp1	Yop
	YscC	mip	Tol
	Pilus	BtuB	

Those ORFs that did not meet the minimum requirement for being an outer membrane protein based on the above search criteria but which were homologous to identified outer membrane ORFs in Chlamydia pneumoniae were included. The Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) was analyzed using the above search criteria and a number of outer membrane ORFs were identified. These Chlamydia pneumoniae ORFs were then tested against the Chlamydia trachomatis genome using Blastp. Any Chlamydia trachomatis ORF with a Blastp P value less than e⁻¹⁰ against a Chlamydia pneumoniae outer membrane was included in this section, if there was no better data indicating otherwise. A list of ORFs in the Chlamydia trachomatis genome encoding putative surface exposed proteins is set forth above in the specification.

Identification of Putative Lipoproteins in the Genome of Chlamydia trachomatis

25 proteins (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108). The characteristic features of lipoproteins are a thiol-linked diacylglyceride and an amine-linked monoacyl group on the cysteine that becomes the amino-terminal residue after signal peptide cleavage by Signal Peptidase II. (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108). The identification of putative lipoproteins from the genomic sequencing of Chlamydia trachomatis was done by examining the deduced amino acid sequence of identified ORFs for the presence of a signal peptide with a Signal Peptidase II cleavage site analogous to the consensus sequence for prolipoprotein modification and processing reactions (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York;

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Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128).

The deduced amino acid sequences of *Chlamydia trachomatis* ORFs were initially screened for the most basic of lipoprotein characteristics, a cysteine in the first 30 amino acids of the deduced protein. ORFs with a standard start codon (ATG, GTG, or TTG) and having one or more of the following characteristics were selected for direct analysis of their first 30 amino acids:

- (a) Significant Signal P value (at least two out-of-the four values are Yes)
- (b) PSORT value indicating membrane passage (IM-inner membrane, Peri-periplasm, or OM-outer membrane)
 - (c) Identification of the word lipoprotein among the ORF Blastp data set.
- (d) A Blastp value of <e⁻¹⁰ with a putative lipoprotein from *Chlamydia pneumoniae* (French application No. 97-14673 filed 21 November 1997).

The first 30 amino acids encoded by each ORF in this set were analyzed for the characteristics commonly found in lipoprotein signal peptides (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108; Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York; Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Grampositive bacteria. J. Bacteriol. 177:1123-1128.) Putative lipoprotein signal peptides were required to have a cysteine between amino acid 10 and 30 and reach a minimum score of three based on the following criteria for lipoprotein signal peptides:

- (a) Identification of specific amino acids in specific positions around the cysteine which are part of the consensus Signal Peptidase II cleavage site (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York); Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128). Since the identification of the cleavage site is the most important factor in identifying putative lipoproteins, each correctly positioned amino acid contributed toward reaching the minimum score of three.
- (b) A hydrophobic region rich in alanine and leucine prior to the cleavage site (Pugsley, A. P., 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.
- (c) A short stretch of hydrophilic amino acids greater than or equal to 1, usually lysine or arginine, following the N-terminal methionine (Pugsley, A. P., 1993. The complete general secretory

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pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.

A list of ORFs in the *Chlamydia trachomatis* genome encoding putative lipoproteins is set forth above in the specification.

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LPS-Related ORFs of Chlamydia trachomatis

Lipopolysaccharide (LPS) is an important major surface antigen of Chlamydia cells. Monoclonal antibodies (Mab) directed against LPS of Chlamydia pneumoniae have been identified that can neutralize the infectivity of Chlamydia pneumoniae both in vitro and in vivo (Peterson et al. 10 1988). Similar results are expected utilizing monoclonal antibodies against LPS of Chlamydia trachomatis. LPS is composed of lipid A and a core oligosaccharide portion and is phenotypically of the rough type (R-LPS) (Lukacova, M., Baumann, M., Brade, L., Mamat, U., Brade, H. 1994. Lipopolysaccharide Smooth-Rough Phase Variation in Bacteria of the Genus Chlamydia. Infect. Immun. June 62(6):2270-2276.) The lipid A component is composed of fatty acids which serve to anchor LPS in the outer membrane. The core component contains sugars and sugar derivatives such as a trisaccharide of 3-deoxy-D-manno-octulosonic acid (KDO) (Reeves, P.R., Hobbs, M., Valvano, M.A., Skurnik, M., Whitfield, C., Coplin, D., Kido, N., Klena, J., Maskell, D., Raetz, C.R.H., Rick, P.D. 1996. Bacterial Polysaccharide Synthesis and Gene Nomenclature pp. 10071-10078, Elsevier Science Ltd.). The KDO gene product is a multifunctional glycosyltransferase and represents a 20 shared epitope among the Chlamydia. For a review of LPS biosynthesis see, e.g., Schnaitman, C.A., Klena, J.D. 1993. Genetics of Lipopolysaccharide Biosynthesis in Enteric Bacteria. Microbiol. Rev. 57:655-682.

A text search of the ORF Blastp results identified several genes that are involved in Chlamydial LPS production with a P score less than e⁻¹⁰. The following key-terms were used in the text search: KDO, CPS (Capsular Polysaccharide Biosynthesis), capsule, LPS, rfa, rfb, rfc, rfe, rha, rhl, core, epimerase, isomerase, transferase, pyrophosphorylase, phosphatase, aldolase, heptose, manno, glucose, lpxB, fibronectin, fibrinogen, fucosyltransferase, lic, lgt, pgm, tolC, rol, ChoP, phosphorylcholine, waaF, PGL-Tb1. A list of ORFs in the Chlamydia trachomatis genome encoding putative polypeptides involved in LPS biosynthesis is set forth above in the specification.

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Type III And Other Secreted Products

Type III secretion enables gram-negative bacteria to secrete and inject pathogenicity proteins into the cytosol of eukaryotic host cells (Hueck, C. J., 1998. Type III Protein Secretion Systems in Bacterial Pathogens of Animals and Plants. In Microbiology and Molecular Biology Reviews. 62:379-433.) These secreted factors often resemble eukaryotic signal transduction factors,

thus enabling the bacterium to redirect host cell functions (Lee, C.A., 1997. Type III secretion systems: machines to deliver bacterial proteins into eukaryotic cells? Trends Microbiol. 5:148-156.) In an attempt to corrupt normal cellular functions, Chlamydial pathogenicity factors injected into the host cytosol will nonetheless, as cytoplasmic constituents be processed and presented in the context of the Major Histocompatibility Complex (MHC class I). As such, these pathogenicity proteins represent MHC class I antigens and will play an important role in cellular immunity. Also included in this set are secreted non-type III products that may play a role as vaccine components.

A text search of the ORF Blastp results identified genes that are involved in *Chlamydia trachomatis* protein secretion with a P score less than e⁻¹⁰. The following key-terms were used in the text search in an effort to identify surface localized or secreted products: Yop, Lcr, Ypk, Exo, Pcr, Pop, Ipa, Vir, Ssp, Spt, Esp, Tir, Hrp, Mxi, hemolysin, toxin, IgA protease, cytolysin, tox, hap, secreted and Mip.

Chlamydia trachomatis ORFs that did not meet the above keyword search criteria, but have homologs in Chlamydia pneumoniae that do meet the search criteria are included herein.

The Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) was analyzed using the above search criteria and a number of ORFs were identified. These Chlamydia pneumoniae ORFs were tested against the Chlamydia trachomatis genome using Blastp. Any Chlamydia trachomatis ORF with a Blastp P value < e⁻¹⁰ against a Chlamydia pneumoniae homolog, identified using the above search criteria, was included. A list of ORFs in the Chlamydia trachomatis genome encoding putative secreted proteins is set forth above in the specification.

Chlamydia trachomatis RGD Recognition Sequence

Proteins that contain Arg-Gly-Asp (RGD) attachment site, together with integrins that serve as their receptor constitute a major recognition system for cell adhesion. The RGD sequence is the cell attachment site of a large number of adhesive extracellular matrix, blood, and cell surface proteins and nearly half of the known integrins recognize this sequence in their adhesion protein ligands. There are many RGD containing microbial proteins such as the penton protein of adenovirus, the coxsackie virus, the foot and mouth virus and pertactin, a 69 kDa (kilodalton) surface protein of Bordetella pertussis, that serve as ligands through which these microbes bind to integrins on the cell surfaces and gain entry into the cell. The following provides evidence supporting the importance of RGD in microbial adhesion:

a) The adenovirus penton base protein has a cell rounding activity and when penton base was expressed in *E. coli*, it caused cell rounding and cells adhered to polystyrene wells coated with the protein. Mutant analysis showed that both these properties required an RGD sequence. Virus mutants with amino acid substitutions in the RGD sequence, showed much less adherence to HeLa S3

cells, and also were delayed in virus reproduction (Bai, M., Harfe, B., and Freimuth, P. 1993. Mutations That Alter an RGD Sequence in the Adenovirus Type 2 Penton Base Protein Abolish Its Cell-Rounding Activity and Delay Virus Reproduction in Flat Cells. J. Virol. 67:5198-5205).

b) It has been shown that attachment and entry of coxsackie virus A9 to GMK cells were dependent on an RGD motif in the capsid protein VP1. VP1 has also been shown to bind ανβ3 integrin, which is a vitronectin receptor (Roivainen, M., Piirainen, L., Hovi, T., Virtanen, I., Riikonen, T., Heino, J., and Hyypia, T. 1994. Entry of Coxsackievirus A9 into Host Cells: Specific Interactions with ανβ3 Integrin, the Vitronectin Receptor Virology, 203:357-65).

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- c) During the course of whooping cough, Bordetella pertussis interacts with alveolar macrophages and other leukocytes on the respiratory epithelium. Whole bacteria adheres by means of two proteins, filamentous hemagglutinin (FHA) and pertussis toxin. FHA interacts with two classes of molecules on macrophages, galactose containing glycoconjugates and the integrin CR3.

 The interaction between CR3 and FHA involves recognition of RGD sequence at the positions 1097-1099 in FHA (Relman, D., Tuomanen, E., Falkow, S., Golenbock, D. T., Saukkonen, K., and Wright, S. D. "Recognitition of a Bacterial Adhesin by an Integrin: Macrophage CR3 Binds Filamentous Hemagglutinin of Bordetella Pertussis." Cell, 61:1375-1382 (1990)).
- d) Pertactin, a 69 kDa outer membrane protein of *Bordetella pertussis*, has been shown to promote attachment of Chinese hamster ovary cells (CHO). This attachment is mediated by recognition of RGD sequence in pertactin by integrins on CHO cells and can be inhibited by synthetic RGD containing peptide homologous to the one present in pertactin (Leininger, E., Roberts, M., Kenimer, J. G., Charles, I. G., Fairweather, N., Novotny, P., and Brennan, M. J. 1991. Pertactin, an Arg-Gly-Asp containing *Bordetella pertussis* surface protein that promotes adherence of mammalian cells Proc. Natl. Acad. Sci. USA, 88:345-349).
- e) The RGD sequence is highly conserved in the VP1 protein of foot and mouth disease virus (FMDV). Attachment of FMDV to baby hamster kidney cells (BHK) has been shown to be mediated by VP1 protein via the RGD sequence. Antibodies against the RGD sequence of VP1 blocked attachment of virus to BHK cells (Fox, G., Parry, N. R., Barnett, P. V., McGinn, B., Rowland, D. J., and Brown, F. 1989. The Cell Attachment Site on Foot-and-Mouth Disease Virus Includes the Amino Acid Sequence RGD (Arginine-Glycine-Aspartic Acid) J. Gen. Virol., 70:625-637).

It has been demonstrated that bacterial adherence can be based on interaction of a bacterial adhesin RGD sequence with an integrin and that bacterial adhesins can have multiple

binding site characteristic of eukaryotic extracellular matrix proteins. RGD recognition is one of the important mechanisms used by microbes to gain entry into eukaryotic cells.

The complete deduced protein sequence of the Chlamydia trachomatis genome was searched for the presence of RGD sequence. There were a total of 38 ORFs that had one or more 5 RGD sequences. Not all RGD containing proteins mediate cell attachment. It has been shown that RGD containing peptides that have proline immediately following the RGD sequence are inactive in cell attachment assays (Pierschbacher & Ruoslahti. 1987. Influence of stereochemistry of the sequence Arg-Gly-Asp-Xaa on binding specificity in cell adhesion. J. Biol. Chem. 262:17294-98). ORFs that had RGD, with proline as the amino acid following the RGD sequence were excluded from 10 the list. Also, RGD sequence may not be available at the surface of the protein or may be present in a context that is not compatible with integrin binding. Since not all RGD-containing proteins are involved in cell attachment, several other criteria were used to refine the list of RGD-containing proteins. A list of ORFs in the Chlamydia trachomatis genome encoding polypeptides with RGD recognition sequence(s) is set forth above in the specification.

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Non-Chlamydia pneumoniae ORFs

Chlamydia trachomatis ORFs were compared to the ORFs in the Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) using Blastp. Any Chlamydia trachomatis ORF with a Blastp «P» value greater than e⁻¹⁰ (i.e. >e⁻¹⁰) against 20 Chlamydia pneumoniae ORFs are included in this section. A list of ORFs in the Chlamydia trachomatis genome which are not found in Chlamydia pneumoniae is set forth above in the specification.

Cell Wall Anchor Surface ORFs

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Many surface proteins are anchored to the cell wall of Gram-positive bacteria via the conserved LPXTG motif (Schneewind, O., Fowler, A., and Faull, K.F. 1995. Structure of the Cell Wall Anchor of Surface Proteins in Staphylococcus aureus. Science 268:103-106). A search of the proteins encoded by the Chlamydia trachomatis ORFs was done using the motif LPXTG. A list of ORFs in the Chlamydia trachomatis genome encoding polypeptides anchored to the cell wall is set 30 forth above in the specification.

ECACC Deposits

Samples of Chlamydia trachomatis were deposited with the European Collection of Cell Cultures (ECACC), Salisbury, Wiltshire SP4 OJG, UK on November 26, 1998 and assigned the 35 provisional accession number 98112618. Cells can be grown, harvested and purified, and DNA can

be prepared as discussed above. In order to enable recovery of specific fragments of the chromosome, one can run targeted PCR reactions, whose amplification products can then be sequenced and/or cloned into any suitable vector, according to standard procedures known to those skilled in the art.

In addition, a pool of clones covering the *Chlamydia trachomatis* genome was deposited with the ECACC on November 26, 1998 and assigned provisional accession number 98112617. The pool of clones contains a series of clones, which when taken together, cover the whole chromosome, with a redundancy of slightly more than ten. The total number of clones in the sample is 13,572.

Table 4 lists groups of oligonucleotides to be used to amplify each of ORFs 2-1197 according to standard procedures known to those skilled in the art. Such oligonucleotides are listed as SEQ ID Nos. 1198 to 5981. For each ORF, the following is listed: one forward primer positioned 2,000 bp upstream of the beginning of the ORF; one forward primer positioned 200 bp upstream of the beginning of the ORF; one reverse primer positioned 2,000 bp downstream at the end of ORF, which is 2,000 bp upstream of the end site of the ORF on the complementary strand; and one reverse primer 200 bp downstream at the end of ORF, which is 200 bp upstream of the end site of the ORF on the complementary strand. The corresponding SEQ ID Nos. for the primers are listed in Table 4, where Fp is the proximal forward primer; Fd is the distal forward primer; Bp is the proximal reverse primer; and Bd is the distal reverse primer. The positions of the 5' ends of each of these primers on the nucleotide sequence of SEQ ID No. 1 are shown in Table 5.

The present invention is not to be limited in scope by the specific embodiments described herein, which are intended as single illustrations of individual aspects of the invention, and functionally equivalent methods and components are within the scope of the invention. Indeed, various modifications of the invention, in addition to those shown and described herein will become apparent to those skilled in the art from the foregoing description and accompanying drawings. Such modifications are intended to fall within the scope of the appended claims.

All publications and patent applications mentioned in this specification are herein incorporated by reference to the same extent as if each individual publication or patent application was specifically and individually indicated to be incorporated by reference.

			TABLE 1				
_	begin	stop	Homology	A	Species	Score	18
Ш	501	208	putative		4		ì
	3276	505	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	379	37
	5068	3242	lipid A disaccharide synthetase (lpxB)	U32786	Haemophilus influenzae	285	40
	6400	5126	poly(A) polymerase	AE000123	Escherichia coli	552	46
	7977	6619	D-alanine permease (dagA)	U32770	Haemophilus influenzae	265	36
<u> </u>	8582	8082	signalpeptidase II	X78084	Staphylococcus carnosus	174	36
	8995	8591	YteA	AF008220	Bacillus subtilis	157	43
	9440	8979	ORF 168	D28752	Synechococcus sp.	318	42
'	9828	10430	unknown	280108	Mycobacterium tuberculosis	324	94
	10367	11254	hypothetical protein (SP:P39587)	U67605	Methanococcus jannaschii	152	38
_ '	11245	11916	rRNA methylase	D90913	Synechocystis sp.	209	6
	12068	13324	hypothetical	U32691	Haemophilus influenzae	367	45
	13532	14413	neutral amino acid transporter B0.	U75284	Oryctolagus cuniculus	410	39
T T	14807	15019	dihydrolipoamide acetyltransferase	L38646	Saccharopolyspora erythraea	324	47
	14932	15969	branched chain alpha-keto acid dehydrogenase E2	M97391	Bacillus subtilis	577	4
	15995	16501	ORF_0328	U18997	Escherichia coli	223	4
	16467	16138	putative				
	18190	17417	putative outer membrane protein	U80956	Borrelia burgdorferi	98	36
	20521	18437	ORF-2	D11024	Shigella flexmeri	642	37
	22202	20814	dnaK like protein (AA 1-660)	X52175	Chlamydia trachomatis	2214	66
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81	8	66	88	_	6	6	6	T	<u></u>					Τ		L	Τ		Τ
-	8	6	6	87	8	66	66	\perp	4	_	igg	45	_	\downarrow	32	4	_	1	_
Score	558	503	580	373	1999	3044	1298		786			126			175	245			
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Synechocystis sp.			Bartonella bacilliformis			Saccharomyces cerevisiae	Caenorhabditis elegans							
A	L22180	M62819	L25105	L25105	125105	L25105	125105		D64004			L46591			U18917	271261			
Homology	ORF, 82 kDa protein	heat shock protein	GrpE-like protein	GrpE-like protein	has homology to putative heat shock proteins of Bacillus subtilis and Clostridium acetobutylicum; ORFA; putative	aminoacyl-tRNA synthetase	ORFB; putative	putative	hypothetical protein	putative	putative	putative	putative	putative	Yer156cp	F21C3.3	putative	putative	putative
stop	22153	22478	22824	23110	23394	24641	27710	27725	28740	30032	30520	31780	31800	33314	35027	35359	35919	36698	37013
begin	22602	22804	23183	23394	24569	26383	26640	28780	29957	30721	31281	31436	33356	33901	34116	34988	35167	35923	37810
ORF	ORF22	ORF23	ORF24	ORF25	Orf26	ORF27	ORF28	ORF29	ORF30	ORF31	ORF32	ORF33	ORF34	ORF35	ORF36	ORF37	ORF38	ORF39	ORF40

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*1	8	4		43	95	9	42				47				36		30	28	38	37			1		4	
Score	529	192		663	681	959	755				892				142	1	462	801	154	355			250.	000	590	
Species	Bacillus subtilis	Escherichia coli		Mycobacterium tuberculosis	Chlamydia trachomatis	Chlamydia trachomatis	Flaveria bidentis				Pseudomonas aeruginosa				Nicotiana tabacum		Synechocystis sn		Escherichia coli	Methanococcus	Jannaschil		Danidonopas arrivas	pv. tabaci	Methanococcus	jannaschii
ΩI	AF008220	X67753		Z98209	L40822	L40822	L40958				M33223				L32794		D90911	X67516	M13045	U67500			114 7017		U67476	
Homology	DAPH synthase-chorismate	arginine binding protein	putative	hypothetical protein MTCY154.05c	phophoglucoisomerase-like protein	phophoglucoisomerase-like protein	NADP-malate dehydrogenase	putative	putative	putative	membrane protein (arcD)	putative	putative	putative	dehydroguinate	dehydratase/shikimate dehydrogenase	3-dehydroquinate synthase	chorismate synthase	shikimate kinase II	. ~	putative	putative	dihydrodipicolinate	reductase	aspartate-semialdehyde	denyarogenase
stop	39085	39927	40756	42007	43116	43802	45227	45275	45975	46506	46882	48178	48630	50099	50925		52348	53466	54595	55031	56084	58235	59181		60195	
begin	38207	39151	39923	40760	42175	42999	44211	46072	46340	46895	47955	48585	50072	20710	52439		53484	54536	55086	56350	55659	56847	58423		59185	
ORF	ORF41	ORF42	ORF43	ORF44	ORF45	ORF46	ORF47	ORF48	ORF49	ORF50	ORF51	ORF52	ORF53	ORF54	ORF55		ORFS6	ORF57	ORF58	ORF59	ORF60	ORF61	ORF62		ORF63	

18	41	42		42				4		æ	43	43	33	7			×		45				38		40	48	
Score	312	345		148				733		156	306	272	6	0			283		1447				495		400	1927	-
Species	Escherichia coli	Helicobacter pylori		Bacillus subtilis				Bacillus subtilis		Mycobacterium tuberculosis	Bacillus subtilis	Calothrix PCC7601	Usemonhilm influence	naciilopiitins tiittueiizad			Bacillus subtilis		Escherichia coli				Haemophilus influenzae		Synechocystis sp.	Arabidopsis thaliana	
A	000000	AE000609		X14084				D26185		Z94752	D26185	X10305	1722727	N			M96343		AE000184				U32750		D64001	U88087	
Homology	aspartokinase III	dihydrodipicolinate synthetase (dapA)	putative	hypothetical protein	putative	putative	putative	unknown	putative	KsgA	high level kasgamycin resistance	polypeptide deformylase	protein translocation	protein, low temperature (secG)	putative	putative	homologous to unidentified B. coli protein	!	, This 530	identical	640 aa procein ihes_hakin SW: P44808	putative	integrase-recombinase	protein (xerC)	hypothetical protein	LON protease homolog	putative
stop	61483	62353	63141	63983	64071	64656	64609	67269	68873	69233	69721	70455	71006		71086	73497	74876		75502			77299	77145		78154	79878	83271
begin	60188	61496	62500	63396	64628	64285	64944	65347	95929	68877	69212	69958	70701		73191	74900	75463		77124			77000	78095		79065	81971	82639
ORF	ORF64	ORF65	ORF66	ORF67	ORF68	ORF69	ORF70	ORF71	ORF72	ORF73	ORF74	ORF75	ORF76		ORF77	ORF78	ORF79		ORF80			ORF81	ORF82		ORF83	ORF84	ORF85

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18	45						32	¥	35		ಜ	36	40	4	42	16	37	49		51	48	æ	
Score	822						128	671	120		542	326	487	2090	319	2411	494	471		634	558	300	
Species	Salmonella typhimurium						Bacillus subtilis	Bacillus stearothermophilus	Bacillus subtilis		Bacillus subtilis	Helicobacter pylori	Helicobacter pylori	Thermus thermophilus	Helicobacter pylori	Chlamydia trachomatis	Synechocystis sp.			Haemophilus influenzae	Bacillus subtilis	Arabidopsis thaliana	
Ð	U58360						X12832	U12340	X17014		X06803	AE000583	AE000583	D49911	AE000583	U83196	D90903	D84432		U32723	L29475	U18969	
Homology	DnaJ	putative	putative	putative	putative	putative	Hpr protein	PTS enzyme I	ORF107	putative	dnaZX-like ORF put. DNA	excinuclease ABC subunit A (uvrA)	excinuclease ABC subunit A (uvrA)	UvrA	excinuclease ABC subunit A (uvrA)	pyruvate kinase	hypothetical protein	YqiE	putative	exonuclease VII, large subunit (xseA)	triose phosphate isomerase	phosphoribosylanthranilate isomerase	putative
stop	84850	86921	87313	87805	88747	89265	89732	91447	91435	91745	92344	93361	94071	94628	98113	98741	100337	101323	102210	102726	104254	105894	108460
pegin	83792	84876	88650	87440	88400	88717	89355	89735	91749	92392	93138	94134	94637	98299	98715	100228	101347	102210	102485	104315	105075	105259	107429
ORF	ORF86	ORF87	ORF88	ORF89	ORF90	ORF91	ORF92	ORF93	ORF94	ORF95	ORF96	ORF97	ORF98	ORF99	ORF100	ORF101	ORF102	ORF103	ORF104	ORF105	ORF106	ORF107	ORF108

ORF	begin	stop	Homology	A	Species	Score	%I
ORF109	108665	108955	putative				
ORF110	109459	109013	putative				
ORF111	110366	109704	putative				
ORF112	111330	112520	elongation factor Tu	L22216	Chlamydia trachomatis	2007	100
ORF113	112915	113463	transcription antitermination protein	U32754	Haemophilus influenzae	313	37
			(nusG)				
ORF114	113566	113994	ribosomal protein L11	D13303	Bacillus subtilis	443	23
ORF115	114020	114604	ribosomal protein L1	Z11839	Thermotoga maritima	528	22
ORF116	114720	115253	ribosomal protein L10	Z11839	Thermotoga maritima	143	88
ORF117	115362	115676	rpl12 (AA 1-128)	X53178	Synechocystis PCC6803	254	62
ORF118	116022	119795	DNA-directed RNA polymerase beta chain	X64172	Staphylococcus aureus	2675	61
ORF119	119823	124010	DNA-directed RNA polymerase beta' chain (rpoC)	U32733	Haemophilus influenzae	3486	20
ORF120	124065	124988	transaldolase	L19437	Homo sapiens	677	20
ORF121	124873	125106	transaldolase	U67611	Homo sapiens	121	4
ORF122	126261	125536	putative				
ORF123	126328	126930	putative				
ORF124	127138	127785	putative				
ORF125	127924	129714	Al isoform of vacuolar H+- ATPase subunit A	U22077	Gallus gallus	1062	45
ORF126	129720	131033	membrane ATPase	X79516	Haloferax volcanii	790	48
ORF127	131018	131629	putative				
ORF128	131834	133156	Na+ -ATPase subunit I	D17462	Enterococcus hirae	188	क्ष
ORF129	133075	133584	v-type Na-ATPase	X76913	Enterococcus hirae	110	38
ORF130	133625	133999	v-type Na-ATPase	X76913	Enterococcus hirae	. 68	32
ORF131	133861	134508	putative				
ORF132	134638	137454	valyl-tRNA synthetase	D64006	Synechocystis sp.	1763	51

1. %	4		38	83	39		4		26	52	83	:	4		43				47	69	45	S	1	39		T
Score	452		282	1113	356		741		625	704	277		168		169				292	555	986	1535		, 66		1
Species	Mycobacterium tuberculosis		Yersinia pestis	Bacillus subtilis	Bacillus subtilis		Streptococcus	pneumoniae	Caenorhabditis elegans	Helicobacter pylori	Mycobacterium	0	Escherichia coli		Escherichia coli				Haemophilus influenzae	Escherichia coli	Bacillus subtilis	Bacillus subtilis		Bacillus subtilis		
Ħ	295209		N22968	D26185	D64116		Z47210		D12984	AE000604	Z96072	2000	1888In		U18997				U32702	AE000213	U02604	U02604		Y08559		
Homology	PknD	putative	porphobilinogen deaminase	unknown	ORF3	putative	unknown		manganese superoxide dismutase precursor	acetyl-CoA carboxylase beta subunit (accD)	Dut	TTANT -	בווקלווופ זושוורן	putative	enzyme IIANtr	putative	putative	putative	hypothetical	hypothetical protein in purB 5' region	ClpC adenosine triphosphatase	ClpC adenosine triphosphatase	putative	Unknown	putative	putative
stop	140276	140335	141077	141780	143128	144393	146326		147078	148075	148549	149027	17061	149305	149708	150911	151004	151999	153352	153997	153984	155231	157525	158955	159961	161220
begin	137442	140733	141799	143240	143829	143923	144548		146413	147140	148115	148524	170011	149000	149187	149712	152044	152664	152900	153389	155276	156544	156806	157489	159104	159916
ORF	ORF133	ORF134	ORF135	ORF136	ORF137	ORF138	ORF139		ORF140	ORF141	ORF142	OPE143	()	ORF144	ORF145	ORF146	ORF147	ORF148	ORF149	ORF150	ORF151	ORF152	ORF153	ORF154	ORF155	ORF156

ORF	pegin	stop	Homology	CI	Species	Score	#1
ORF157	161183	161593	glycine cleavage protein homolog	U12980	Saccharomyces cerevisiae	175	35
ORF158	162354	161623	unidentified protein of Na+- translocating NADH-quinone reductase	D49364	Vibrio alginolyticus	524	51
ORF159	163013	162363	NADH:uniquinone oxidoreductase	Z37111	Vibrio alginolyticus	543	55
ORF160	163941	162994	NADH:ubiquinone oxidoreductase (GP:Z37111_4)	U32702	Haemophilus influenzae	287	22
ORF161	165505	164474	NADH:ubiquinone oxidoreductase subunit B	237111	Vibrio alginolyticus	449	45
ORF162	166686	166093	H. pylori predicted coding region HP1542	AE000652	Helicobacter pylori	111	eg E
ORF163	168171	166729	pot. ORF 446 (aa 1-446)	X02369	Bacillus subtilis	722	42
ORF164	169249	168848	putative				
ORF165	169586	170431	hypothetical protein	D90906	Synechocystis sp.	462	48
ORF166	170780	171334	putative				
ORF167	171333	172376		M26645	Neisseria flavescens	210	47
ORF168	172309	172722	penicillin-binding protein 2	M26645	Neisseria flavescens	176	44
ORF169	173048	174496	murE gene product	Z15056	Bacillus subtilis	789	43
ORF170	174399	174968	N-acetylmuramoyl-L-alanine amidase (amiA)	AE000589	Helicobacter pylori	177	14
ORF171	175267	175710	integration host factor beta subunit	L35259	Pseudomonas aeruginosa	110	88
ORF172	175714	177009	putative				
ORF173	177423	178115	carboxyltransferase alpha subunit	U59236	Synechococcus PCC7942	558	25
ORF174	178084	180021	ATP dependent translocator homolog (msbA)	U32691	Haemophilus influenzae	453	4
ORF175	180704	180048	putative				

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1%	쫎	S	ಜ	43	30	જ્		98		æ				40	£3		26	ಜ	42	S
Score	256	173	371	452	93	154		96		66				373	545		1758	580	148	795
Species	Helicobacter pylori	Arabidopsis thaliana	Bacillus subtilis	Bacillus subtilis	Escherichia coli	Haemophilus influenzae		Symechocystis sp.		Rhodobacter sphaeroides				Helicobacter pylori	Staphylococcus aureus		Homo sapiens	Escherichia coli	Saccharomyces cerevisiae	Caenorhabditis elegans
αī	AE000536	AF007270	Y13937	AF008220	D90888	U32728		D90902		AJ000977				AE000645	D89066		U47025	X16931	X86470	277659
Homology	H. pylori predicted coding region HP0152	contains similarity to DNA polymerase III, alpha chain (SP:P47277)	putative Ptcl protein	NifS2	similar to [SwissProt Accession Number P37908]	hypothetical	putative	regulatory protein for beta- lactamase	putative	prolipoprotein diacylglyceryl transferase	putative	putative	putative	60 kDa inner-membrane protein	DnaA	putative	glycogen phosphorylase B	glycogen phosphorylase (AA 1 - 790)	unknown	F23B12.5
stop	180631	181398	183656	184786	184796	186000	186749	187809	188798	190352	190510	191786	192464	193183	194630	194690	197031	197635	198208	197668
pegin	181398	182594	182895	183665	186007	186848	187270	187426	189481	189693	190235	190785	191790	192392	193254	195046	195184	197018	197762	198963
ORF	ORF176	ORF177	ORF178	ORF179	ORF180	ORF181	ORF182	ORF183	ORF184	ORF185	ORF186	ORF187	ORF188	ORF189	ORF190	ORF191	ORF192	ORF193	ORF194	ORF195

1%	48	31	99	43		æ			31	40	92	48	51	22	31	T	83	43
Score	856	170	302	127		577			83	334	706	522	919	220	150		852	231
Species	Arabidopsis thaliana	Porphyra purpurea	Thiobacillus ferrooxidans	Clostridium magnum		Escherichia coli			Brucella abortus	Synechocystis sp.	Bscherichia coli	Bacillus subtilis	Bacillus subtilis	Haemophilus influenzae	Rattus norvegicus		Chlamydia psittaci	Chlamydia psittaci
OI.	U09137	U38804	U81808	L31844		U70214			U51683	D90916	M77744	U59433	U59433	U32701	J02934		AF017105	AF017105
Homology	pyruvate dehydrogenase El beta subunit	pyruvate dehydrogenase El component, alpha subunit	pyruvate dehydrogenase complex Bl alpha subunit	TPP-dependent acetoin dehydrogenase alpha-subunit	putative	UDP-3-0-[3-hydroxymyristoy1]	gracosamine n- acyltransferase	putative	OMP1 precursor	recombination protein	beta-ketoacyl-acyl carrier protein synthase III	malonyl-CoA:Acyl carrier protein transacylase	3-ketoacyl-acyl carrier protein reductase	acyl carrier protein (acpP)	protein kinase type II regulatory subunit (, EC 2.7.1.37)	putative	unknown	inclusion membrane protein C
stop	198962	199941	200266	200585	202377	202380		203471	204059	206811	208528	209471	210214	210816	210883	211374	212875	214168
pegin	199957	200327	200685	200962	201169	203441		203998	206449	207425	207506	208545	209471	210586	211332	212978	214134	214710
ORF	ORF196	ORF197	ORF198	ORF199	ORF200	ORF201		ORF202	ORF203	ORF204	ORF205	ORF206	ORF207	ORF208	ORF209	ORF210	ORF211	ORF212

Г	1	T	T	Т	Т	7	$\overline{}$	T	Т-	1	Τ-	1	1	_	7			Т-	_			 	
18	47	8	8										66	46	25		45	98		48	36		35
Score	181	1341	1027										619	530	334		96	321		274	246	•	740
Species	Chlamydia psittaci	Chlamydia psittaci	Chlamydia psittaci										Chlamydia trachomatis	Bacillus subtilis	Methanococcus	jannaschii	Saccharomyces cerevisiae	Bacillus firmus		Helicobacter pylori	Escherichia coli		Lactococcus lactis
ΕĪ	AF017105	AF017105	AF017105										U57090	D84432	U67467		U43834	U61168		AE000602	AE000232	 11	AF005098
Homology	inclusion membrane protein B	sodium-dependent transporter	amino acid transporter	putative	putative	putative	putative	putative	putative	putative	putative	putative	LAGLI-DADG endonuclease	YqfU	phenylacrylic acid	qecarboxy.lase	Ydr537cp	4-hydroxybenzoate octaprenyltransferase	putative	stationary-phase survival protein (surB)	f311; This 311 aa ORF is 22 pct identical (13 gaps) to	488 aa protein YACA_BACSU SW: P37563; pyul of D21139	GadC
stop	214754	215236	216892	217441	218702	219009	219748	220430	221074	221541	222092	223290	223818	225171	225174		225549	225749	226769	227161	227750		228607
begin	215143	216705	217917	217088	218364	218695	219179	168612	220499	221137	221601	222472	223423	224278	225749		225334	226654	227299	227646	228457		230001
ORF	ORF213	ORF214	ORF215	ORF216	ORF217	ORF218	ORF219	ORF220	ORF221	ORF222	ORF223	ORF224	ORF225	ORF226	ORF227		ORF228	ORF229	ORF230	ORF231	ORF232		ORF233

re I%	46	35	+-			5 58	3 45	37	35	47	66	86	100	91	100
Score	601	103	482	-	-	1925	1963	307	162	218	914	1770	403	320	2828
Species	Bacillus subtilis	Synechocystis sp.				Salmonella typhimurium	Aeromonas salmonicida	Bacillus subtilis	Haemophilus influenzae	Anabaena sp.	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis
a	D88802	D90905	L33777			X07916	L47978	D26185	U32802	U14553	U83195	U83195	U15192	U15192	U15192
Homology	P. haemolytica o- sialoglycoprotein endopeptidase; P36175 (660) transmembrane	Mg2+ transporter	tRNA guanine transglycosylase	putative	putative	subunit B of DNA gyrase	DNA gyrase	unknown	replication protein (dnaX)	putative isozyme of glucose- 6-P-dehydrogenase; developmentally regulated gene in heterocyst development	glucose 6-phosphate dehydrogenase	glucose 6-phosphate dehydrogenase	ORF3	ORF3	CTP synthetase
stop	252066	256718	257844	258690	259187	261604	264129	264742	265628	265631	266426	266942	268066	268205	268500
pegin	253088	255153	256762	257911	258780	259193	261622	264125	264741	266416	266938	267961	268320	268510	270116
OKK	ORF254	ORF255	ORF256	ORF257	ORF258	ORF259	ORF260	ORF261	ORF262	ORF263	ORF264	ORF265	ORF266	ORF267	ORF268

ORF	pegin	stop	Homology	a	Species	grore	12
ORF269	270892	270095	CMP-2-keto-3-	U15192	Chlamvdia trachomatis	1313	100
			deoxyoctulosonic acid				}
			synthetase				
ORF270	271191	271613	putative				
ORF271	272219	272932	nitrate transporter	X61625	Synechococcus sp.	300	श्र
ORF272	272884	273588	putative				
ORF273	274816	273596	putative				
ORF274	274821	275666	putative				
ORF275	277689	276103	ORF_f535	U28377	Escherichia coli	396	38
ORF276	278268	278816	putative		1		
ORF277	279771	279013	tryptophan synthase alpha subunit	M15826	Pseudomonas aeruginosa	357	37
ORF278	280777	279767	tryptophan synthetase	M91661	Coprinus cinereus	1042	85
ORF279	281603	281295	tryptophan repressor	L26582	Enterobacter aerogenes	151	35
ORF280	282104	281787	putative				
ORF281	284335	282794	putative				
ORF282	284460	284795	putative				
ORF283	284817	285674	putative				
ORF284	285637	286137	putative				
ORF285	286357	286677	putative				
ORF286	286681	287898	hypothetical protein	U88070	Chlamydia psittaci	66	35
ORF287	288127	289227	COME ORF1	D64002	Synechocystis sp.	90	46
ORF288	289744	290679	hypothetical protein	U88070	Chlamydia psittaci	246	36
ORF289	290828	291535	putative				
ORF290	291514	292230	endonuclease	U09868	Escherichia coli	160	37
ORF291	292326	293048	putative				
ORF292	293330	294853	putative				T
ORF293	295684	295010	glutamine transport ATP- binding protein Q	U67524	Methanococcus jannaschii	407	æ
ORF294	296336	295692	H. influenzae predicted coding region HI1555	U32830	Haemophilus influenzae	134	37
ORF295	297238	296243	putative				

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H-		40			41	40	37	25				98	37		45	41	88			
Score		82			745	197	292	316				123	202		169	147	167			
Species		Escherichia coli			Thermus aquaticus	Escherichia coli	Mycoplasma pneumoniae	Mycobacterium tuberculosis				Escherichia coli	Bacillus subtilis		Klebsiella pneumoniae	Bacillus subtilis	Spirulina platensis			
Ωī		U73857			M74792	D90870	AE000047	Z84395				D90832	M57689		X58778	238002	870419			
Homology	putative	similar to putative		putative	DNA ligase	DNA LIGASE (EC 6.5.1.2) (POLYDEOXYRIBONUCLEOTIDE SYNTHASE (NAD+)).	Mycoplasma pneumoniae, DNA ligase; similar to Swiss- Prot Accession Number P15042, from E. coli	ипкломп	putative	putative	putative	Preprotein translocase SecA subunit.	sporulation protein	putative	orfX gene product	Similar to Saccharomyces cerevisiae SUAS protein	serine esterase [Spirulina platensis, C1, Peptide, 207 aa]	putative	putative	putative
stop	298735	300458	300527	302071	304973	306111	306436	306977	309276	309872	310716	311972	314364	314738	314741	315665	316529	317338	317499	317874
begin	297791	298905	302152	304917	306157	306494	306963	308773	309881	310720	311570	312451	313435	314340	315526	316507	317284	317592	318470	317599
ORF	ORF296	ORF297	ORF298	ORF299	ORF300	ORF301	ORF302	ORF303	ORF304	ORF305	ORF306	ORF307	ORF308	ORF309	ORF310	ORF311	ORF312	ORF313	ORF314	ORF315

*		9	3					73	3	9	\$	55	3	8		26	28	52	8	3.5	3					T		T		23
Score		802	200					430	343		3/1	315	1	269		297	1089	208	297	483					1				1	2005
Species		Chlamydia psittaci	1					Bacillus subtilis	Lactobacillus fermentum	St.mococodoents	Jucanococcus ap.	Mycobacterium	tuberculosis	Escherichia coli		Bacillus subtilis	Anabaena sp.	1			ı									Bscherichia coli
A		L35036						D84432	U97348	AR000111		Z95389		U18997		D84432	L14862	D84432	D64001	Y13937										M29364
Homology	putative	ORF2	putative	putative	putative	putative	putative	Yqiz	integral membrane protein homolog	adenylate kinase	putative	RpsI		50S ribosomal subunit	protein L13	хүбх	biotin carboxylase	Хфһ	elongation factor P	putative CfxE protein	putative	ATP-dependent protease binding subunit								
stop	318477	320142	321497	321937	322362	323140	324913	324977	325621	327203	328150	328204		328734		329292	329608	330670	331177	331721	333021	333589	333806	334361	334729	335602	335194	335903	336338	337347
begin	318947	319342	320544	321485	321901	322301	323144	325621	326268	326469	327281	328605		329066		329663	330666	331161	331731	332404	332779	333005	334357	334089	335142	335195	335673	336334	337378	339947
ORF	ORF316	ORF317	ORF318	ORF319	ORF320	ORF321	ORF322	ORF323	ORF324	ORF325	ORF326	ORF327		ORF328		ORF329	ORF330	ORF331	ORF332	ORF333	ORF334	ORF335	ORF336	ORF337	ORF338	ORF339	ORF340	ORF341	ORF342	ORF343

341847 Pz-peptidase
drong
342470 hypa protein
343370 heat shock protein
344032 hypB protein
344225 hypB protein
345142 orf 3'of chaperonin homol hypB [Chlamydia psittaci, pigeon strain P-1041, Peptide Partial, 98 aa]
o247; This 247 aa ORF is 5 pct identical (0 gaps) to 117 residues of an approx. 160 aa protein YPH7_CHRVI SW: P45371
mutY homolog
hypothetical 36.0 kD p in rne-rpmF intergenic region
putative
enoyl-acyl carrier protein reductase [Brassica napus, Peptide, 385 aa]
hypothetical protein
putative

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ă.			8	8	4	100	100	88	32	42	49	25	88	93	87	37	29	62	46	89	49	48	T
Score		213	577	417	1305	948	1216	3311	116	362	192	978	1631	516	2817	585	528	1362	182	1926	286	379	
Species	Do 11 12	bacillus subtilis	Arabidopsis thaliana	Neurospora crassa	Escherichia coli	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Bacillus subtilis	Synechocystis sp.	Pseudomonas fluorescens	Escherichia coli	Chlamydia psittaci		Chlamydia psittaci	Chlamydia psittaci	Chlamydia psittaci	Chlamydia psittaci	Haemophilus influenzae	Chlamydia psittaci	1		
A	70001400	A5001488	Z23108	D45049	D90729	U74759	U74759	U74759	218631	D90917	M35367	AE000219	U88070	U88070	U88070	U88070	U88070	U88070	U32776	U88070	U88070	U88070	
Homology	FUNCTION INKNOWN SIMILAD	IN E. COLI	NADPH thioredoxin reductase	Thioredoxin Reductase (NADPH)	30S ribosomal protein S1	NusA	NusA		ORF6 gene product	tRNA pseudouridine 55 synthase	protein X	hypothetical GTP-binding protein in pth 3' region	cds1 gene product	cds2 gene product	gene	gene	il gene prod		ribosomal protein L28 (rpL28)	hypothetical protein	hypothetical protein	hypothetical protein	putative
stop	353305		353670	354140	356672	357377	358093	360743	361121	361884	362746	362816	362195	365587	367320	368603	369081	370251	371086	372816	373529	374204	374224
pegin	353709		354218	354721	354966	356700	357326	358035	360753	361162	361826	363859	364116	365198	365479	367341	368644	369088	370769	371203	373119	373614	374736
ORF	ORF358		ORF359	ORF360	ORF361	ORF362	ORF363	ORF364	ORF365	ORF366	ORF367	ORF368	ORF369	ORF370	ORF371	ORF372	ORF373	ORF374	ORF375	ORF376	ORF377	ORF378	ORF379

1%	Γ	80			10	T		T	Τ	T			Γ	Τ	Τ.	Τ	Τ	T	Τ	Τ	Τ	T			T
-	L	88	5	_	45	51	86	35		35	-	45	4		32	42	5	31	51	8	_	8		8	22
Score		490	678		137	216	92	477		134		185	989		150	168	134	164	592	282		1295		1235	543
Species		Chlamydia trachomatis	Synechocystis sp.		Haemophilus influenzae	Synechocystis sp.	Haemophilus influenzae	Bacillus subtilis		Escherichia coli		Helicobacter pylori	Synechocystis sp.	4	Streptococcus parasanguis	Bacillus subtilis	Treponema pallidum	Cyanophora paradoxa	Treponema pallidum	Bacillus subtilis		Arabidopsis thaliana		Bacillus subtilis	Helicobacter pylori
a		L40838	D64000		U32702	D90914	U32780	D26185		U70214		AE000606	D64000		M26130	AF008220	U55214	U30821	U55214	AF008220		Z49227		X91655	AE000552
Homology	putative	corresponds to a 97 amino acid long polypeptide	methylenetetrahydrofolate dehydrogenase	putative	hypothetical	small protein	DNA polymerase III beta- subunit (dnaN)	recombination protein	putative	hypothetical	putative	conserved hypothetical secreted protein	hypothetical protein	putative	ORF1; putative	ytgD	TroR	putative protein of 299 amino acids	TroB	YtgA	putative	adenine nucleotide	locae	lepA gene product	GTP-binding membrane protein (lepA)
stop	374703	376748	378737	379048	379403	379641	381470	382567	383702	383655	383949	385178	385706	386627	386872	387338	387816	388237	389173	389945	391810	393996		395354	395992
pegin	376391	377062	377853	378626	379017	380009	380187	381473	382704	383945	385217	385507	386845	386127	387372	387823	388250	389169	389955	390988	391514	392410		394170	395309
ORF	ORF380	ORF381	ORF382	ORF383	ORF384	ORF385	ORF386	ORF387	ORF388	ORF389	ORF390	ORF391	ORF392	ORF393	ORF394	ORF395	ORF396	ORF397	ORF398	ORF399	ORF400	ORF401	00,1100	ORF402	ORF403

4 396538 396059 phosphogluconate dehydrogenase dehydrogenase 6-phosphogluconate dehydrogenase 398753 397401 tyrosyl-tRNA synthetase 5 398753 397401 tyrosyl-tRNA synthetase 6 400167 39978 FLHA gene product 7 401224 400034 flbF flbF flbF flbF flbF flbF flbF flbF	•	SDecies	Score	æ H
397507 396542 6-phosphogluconate dehydrogenase dehydrogenase 398753 397401 Lyrosyl-tRNA synthetase 399688 398909 whiG-Stv gene product 400167 399778 FLHA gene product 401224 400034 flbF flbF ferredoxin IV 1 1 1 1 1 1 1 1 1	U30255	Homo sapiens	411	જ
5 398753 397401 tyrosyl-tRNA synthetase 7 399688 398909 whiG-Stv gene product 8 400167 399778 FLHA gene product 9 401224 400034 flbF 1 401776 402021 ferredoxin IV 1 402126 403220 putative 1 40348 405180 GcpB 2 40378 405520 YfiH 3 405165 405925 dihydrolipoamide 4 405165 405955 dihydrolipoamide 4 405165 405955 dihydrolipoamide 4 405165 405955 dihydrolipoamide 4 405955 dihydrolipoamide dehydrogenase 407049 405955 dihydrolipoamide dehydrogenase 410532 411416 YqeR 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413341 413978 putative 414379 413978	AB006102	Candida albicans	908	51
399688 398909 whiG-Stv gene product 400167 399778 FLHA gene product 401224 400034 flbF 401224 400034 flbF 40176 402021 ferredoxin IV 402126 403220 putative 403348 405180 GcpE 403788 405202 YfiH 405165 40595 dihydrolipoamide 1 407049 405955 dihydrolipoamide 2 2.3.1.61 2.3.1.61 409773 407056 alpha-ketoglutarate 411070 411707 413410 putative 413404 413404 413952 putative 413404 413404 413952 putative 414379 41664 415177 putative 416664 417456 415777 putative	M13148	Bacillus caldotenax	844	45
40167 399778 FLHA gene product 401224 400034 flbF 401776 402021 ferredoxin IV 402126 403220 putative 403348 405180 GcpE 4033788 405920 YfiH 407049 405955 dihydrolipoamide 407049 405955 dihydrolipoamide 409773 407056 alpha-ketoglutarate 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 4134379 413978 putative 414379 putative 416664 415177 putative 416664 415177 putative 417456 416740 unknown	X68709	Streptoverticillium	463	41
401224 400034 flbF 401776 402021 ferredoxin IV 402126 403220 putative 403348 405180 GcpE 403788 403276 putative 405165 405920 YfiH 407049 405955 dihydrolipoamide transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413439 413978 putative 416664 415177 putative 416664 415177 putative 417456 416740 unknown	X63698	Bacillus subtilis	134	35
402126 403220 putative 402126 403220 putative 403348 405180 GcpE 403788 405920 YfiH 405165 405920 YfiH 407049 405955 dihydrolipoamide Lranssuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413841 415112 putative 414379 413978 putative 415577 putative 416664 415177 putative 417456 416740 unknown	M73782	Caulobacter crescentus	355	39
402126 403220 putative 403348 405180 GcpE 403788 405226 YfiH 407049 405955 dihydrolipoamide 407049 405955 dihydrolipoamide 407049 405955 dihydrolipoamide 407073 407056 alpha-ketoglutarate 410532 411416 YqeR 411707 413410 putative 413404 413952 putative 411841 415112 putative 411864 415177 putative 411756 416740 unknown	M59855	Rhodobacter capsulatus	98	22
403348 405180 GCpE 403788 403276 putative 405165 405920 YfiH 407049 405955 dihydrolipoamide transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate dehydrogenase 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413404 413952 putative 414379 413978 putative 41664 415177 putative 417456 416740 unknown				
405165 405920 YfiH 407049 405955 dihydrolipoamide 407049 405955 dihydrolipoamide transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate dehydrogenase 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413404 413952 putative 414379 413978 putative 416664 415177 putative	D90908	Synechocystis sp.	995	49
405165 405920 YfiH 407049 405955 dihydrolipoamide transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate dehydrogenase 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413841 415112 putative 414379 415177 putative 416664 415177 putative 417456 416740 unknown				
407049 405955 dihydrolipoamide transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate dehydrogenase 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413404 413952 putative 414379 413978 putative 414379 415177 putative 417456 416740 unknown	U50134	Escherichia coli	166	43
transsuccinylase (odhB; BC 2.3.1.61) 409773 407056 alpha-ketoglutarate dehydrogenase 411416 YqeR 411707 413410 putative 413434 412606 putative 413404 413952 putative 414379 413978 putative 414379 415177 putative 415177 putative 417456 416740 unknown	M27141	Bacillus subtilis	833	61
409773 407056 alpha-ketoglutarate dehydrogenase 410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413404 413952 putative 413841 415112 putative 414379 413978 putative 41664 415177 putative 417456 416740 unknown				***************************************
410532 411416 YqeR 411707 413410 putative 413433 412606 putative 413404 413952 putative 413841 415112 putative 414379 413978 putative 416664 415177 putative 417456 416740 unknown	U41762	Rhodobacter capsulatus	1537	æ
411707 413410 putative 413433 412606 putative 413404 413952 putative 413841 415112 putative 414379 413978 putative 416664 415177 putative 417456 416740 unknown	D84432	Bacillus subtilis	496	4
413433 412606 putative 413404 413952 putative 413841 415112 putative 414379 413978 putative 416664 415177 putative 417456 416740 unknown				
413404 413952 putative 413841 415112 putative 414379 413978 putative 416664 415177 putative 417456 416740 unknown				
413841 415112 putative 414379 413978 putative 416664 415177 putative 417456 416740 unknown				
414379 413978 putative 416664 415177 putative 417456 416740 unknown				T
416664 415177 putative 417456 416740 unknown				
417456 416740 unknown				
	294752	Mycobacterium tuberculosis	172	မ္က
ORF425 418053 417721 putative				T
ORF426 418603 418031 putative				

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at a	 	96	14	38		55	1	53	8	3		88				53	¥		47	47	26		22
Score	1661	612	269	165		1229		633	613	770		173				289	817		395	287	374		199
Species	Chlamydia trachomatis	Chlamydia trachomatis	Mycoplasma salivarium	Mycobacterium leprae		Synechocystis sp.		Pseudomonas aeruginosa	Helicohanter puloui	110174		Saccharomyces cerevisiae				Escherichia coli	Schizosaccharomyces pombe		Lactococcus lactis	Synechocystis sp.	1 17		Haemophilus influenzae
A	L10193	M86605	D17450	L39923		D90908		D83138	AROOOS54			U27182				U29581	AB004537		AF005098	X72627	U32705		U32705
Homology	Hc2 nucleoprotein	[karp] gene products	aminopeptidase	putative	putative	glycogen operon protein GlgX	putative	Holliday junction specific DNA helicase	deoxycytidine triphosphate	deaminase (dcd)	putative	biotin apo-protein ligase	putative	putative	putative	exonuclease V alpha-subunit	methionyl-tRNA synthetase	putative	RNAseH II	ribosomal protein L19	ł.	methyltransferase (trmD)	tRNA (guanine-N1)- methyltransferase (trmD)
stop	418647	419672	420245	421518	423043	425079	425146	426245	427817		429886	429857	430323	431787	431987	434475	434620	436272	436567	437894	438285		438986
pegin	419531	420190	421171	421988	422486	423226	426054	426985	427248		429560	430360	430637	430933	431658	432232	436308	436574	437685	438262	439127		439339
ORF	ORF427	ORF428	ORF429	ORF430	ORF431	ORF432	ORF433	ORF434	ORF435		ORF436	ORF437	ORF438	ORF439	ORF440	ORF441	ORF442	ORF443	ORF444	ORF445	ORF446		ORF447

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å.	39	40	37	62	4	69	55					25	31	88	25	42		31	88	51				NA A
Score	168	865	314	708	268	704	1687					820	105	388	537	324		77	114	148				892
Species	Haemophilus influenzae	Escherichia coli	Bacillus subtilis	Haemophilus influenzae	Synechocystis sp.	1	Homo sapiens					Bacillus subtilis	Mycobacterium tuberculosis	Pseudomonas aeruginosa	Pseudomonas aeruginosa	Bacillus subtilis		Escherichia coli	Bacillus subtilis	Bacillus subtilis				Synechocystis sp.
αI	U32705	AE000347	249782	U32830	D90904	U04953	U04953					297025	295554	Y10528	Y10528	AB002150		X61000	U87792	275208				D90913
Homology	ribosomal protein S16 (rpS16)	signal recognition particle protein	product similar to E.coli PRFA2 protein	polypeptide chain release factor 1 (prfA)	leader peptidase I	isoleucyl-tRNA synthetase	isoleucyl-tRNA synthetase	putative	putative	putative	putative	product similar to E. coli PhoH protein	CydB	cyanide insensitive terminal oxidase	cyanide insensitive terminal oxidase	Ybbp	putative	HtrB protein	unknown	hypothetical protein	putative	putative	putative	PET112
stop	439358	439699	441042	441898	446388	446452	447932	451076	451144	451517	452632	454868	454972	455367	456047	458450	459632	461203	461196	462621	462895	464629	465848	466113
pegin	439705	441042	441911	442593	444505	448068	449575	450546	451623	452593	453195	453567	455430	456047	457384	457659	458508	459839	461624	461887	463758	464048	464721	467420
ORF	ORF448	ORF449	ORF450	ORF451	ORF452	ORF453	ORF454	ORF455	ORF456	ORF457	ORF458	ORF459	ORF460	ORF461	ORF462	ORF463	ORF464	ORF465	ORF466	ORF467	ORF468	ORF469	ORF470	ORF471

3 I &	46	$oxed{L}$				33	}	39	88			88	9		35		9			88			4	स्र	 39	•					25
Score	1051					173) 	175	193			100	537		234		313			391			114	77	506				-		1624
Species	Moraxella catarrhalis					Chlamydia psittaci	1	Chlamydia psittaci				Chlamydia psittaci	Chlamydia psittaci		Chlamydia psittaci		Chlamydia psittaci			Chlamydia psittaci			Chlamydia psittaci	Chlamydia psittaci	Chlamydia psittaci						Synechococcius PCC6301
ΩI	U49269					U72499		U65942	U72499			U65942	U72499		U72499		U72499			U72499			U65942	U72499	U72499						M31544
Homology	amidase	putative	putative	putative	putative	putative 98 kDa outer	membrane protein	POMP90A precursor	putative 98 kDa outer	membrane protein	putative	POMP91A		membrane protein	putative 98 kDa outer	membrane protein	putative outer membrane	protein	putative	putative 98 kDa outer	membrane protein	putative	POMP90A precursor			membrane protein	putative	putative	putative	putative	branching enzyme
stop	467419	468906	469675	469826	471106	473267		473695	474527		474602	475613	476517		478665		479088		479668	479895		481496	483429	484964	487864		485222	489247	488233	490456	490507
begin	468891	469280	469349	471226	471624	471954		473252	473982		475198	476527	478640		479084		479723		480012	481466		481732	481864	483402	484898		485725	488204	488571	489440	492765
ORF	ORF472	ORF473	ORF474	ORF475	ORF476	ORF477		ORF478	ORF479		ORF480	ORF481	ORF482		ORF483		ORF484		ORF485	ORF486		ORF487	ORF488	ORF489	ORF490		ORF491	ORF492	ORF493	ORF494	ORF495

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14°	\downarrow	_	4	84		\perp	_		\perp			20	\perp		42		4		2	\$	57	31			88		46			
Score			230	245								702			102		740		117	770	386	82			208		384	-		
Species			Bacillus subtilis	Escherichia coli								Haemophilus influenzae			Mycobacterium	tuberculosis	Escherichia coli		Myxococciie yanthiie		Synechocystis sp.				Methanococcus	Jamaschii	Escherichia coli			
Ð			D84432	M54884								U32781			296072		AE000219		AF013216		D64000	U32720			U67608		M15106			
Homology	putative	putative	УqкM	xprB	putative	penicillin tolerance protein (lytB)	putative	putative	hypothetical protein		hypothetical protein in pth- prs interdenic region		fumarate hydratase		fumarase	thiamine-repressed protein	(nmt1)	putative	hypothetical protein	,	methionine amino peptidase	putative	putative	putative						
stop	492893	492737	494675	494869	495365	494872	496634	497176	498515	499239	500732	062005	501808	502692	503722		506986	507439	507649	508590	508478	510691		511527	512185		513092	513055	515244	515269
begin	492357	493744	493875	494573	494835	495174	495687	496295	497703	498280	499215	501710	502863	503675	505002		505739	506999	508404	508291	508915	209600		511039	511547		512382	514287	514789	514994
ORF	ORF496	ORF497	ORF498	ORF499	ORF500	ORF501	ORF502	ORF503	ORF504	ORF505	ORF506	ORF507	ORF508	ORF509	ORF510		ORF511	ORF512	ORF513	ORF514	ORF515	ORF516		ORF517	ORF518	0.550	OKFS19	ORF520	ORF521	ORF522

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#H		L		51	52	64	47	22	49	76		4							52	9	47	જ	43
Score				340	245	306	387	860	970	314		148							476	164	230	452,	488
Species				Porphyromonas gingivalis	Synechocystis sp.	1	Spinacia oleracea	Escherichia coli	Escherichia coli	Haemophilus influenzae		Porphyra purpurea							Bacillus subtilis	Bacillus subtilis	Bacillus stearothermophilus	Mycobacterium tuberculosis	Synechococcus PCC7942
a				X95938	D90901	AF010578	U52048	U18997	U01376	U32812		U38804							X53057	275208	X16188	Z85982	U86147
Homology	putative	putative	putative	orf150 gene product	30S ribosomal protein S15	polynucleotide phosphorylase	polyribonucleotide phophorylase	polynucleotide phosphorylase	ATP-binding protein	cell division protein (ftsH)	putative	ORF327 gene product	putative	putative	putative	putative	putative	putative	phenylalanyl-tRNA synthetase alpha subunit	phenylalany-tRNA synthetase beta subunit	ribosomal protein L20 (AA 1-119)	unknown	UDP-N- acetylenolpyruvylglucosamine reductase
stop	515804	516422	517171	517400	518380	518822	519516	520497	520718	521886	522143	523623	525746	526078	526400	526735	526851	528292	529142	529624	530223	530737	533272
begin	515553	515808	516476	517927	518096	518403	518923	519577	521986	522131	523495	524591	524652	525731	525939	526301	528323	528861	529723	530166	530543	531378	532370
ORF	ORF523	ORF524	ORF525	ORF526	ORF527	ORF528	ORF529	ORF530	ORF531	ORF532	ORF533	ORF534	ORF535	ORF536	ORF537	ORF538	ORF539	ORF540	ORF541	ORF542	ORF543	ORF544	ORF545

18	88	35	33	51	49		39	84		46	88	94	83		જ	46	46	T
Score	273	170	397	1447	226		82	130		221	153	981	869		388	498	330	1
Species	Bacillus subtilis	Mycobacterium tuberculosis	Helicobacter pylori	Helicobacter pylori	Helicobacter pylori		Escherichia coli	Saccharomyces cerevisiae		Clostridium perfringens	Saccharomyces cerevisiae	Helicobacter pylori	Dictyostelium discoideum		Bacillus subtilis	Thermus aquaticus flavus	Helicobacter pylori	
t t	AF008220	292669	AE000553	AE000581	AE000614		AE000459	U33007		X86493	X70951	AE000610	U23408		AJ000975	X54073	AE000588	
Homology	YtqB	hypothetical protein MTCY08D5.03c	ribonucleoside diphosphate reductase, beta subunit (nrdB)	ribonucleoside-diphosphate reductase l alpha subunit (nrdA)	phosphatidylserine synthase (pssA)	putative	hypothetical 54.7 kD protein in udp 3' region precursor (0475)	Ydr430cp; CAI: 0.15	putative	hypA gene product	orfl gene product	serine protease (htrA)	succinyl coenzyme A synthetase alpha subunit	putative	putative succinyl-coA synthetase beta chain	succinateCoA ligase (ADP-forming)	cell division protein (ftsY)	putativa
stop	533244	533944	534878	535956	540519	540969	541805	541825	543222	544179	544487	544951	546584	547382	547476	547900	549459	549663
begin	533849	534672	535915	539153	187983	540523	540906	543255	544133	544565	544762	546423	547480	546789	547901	548634	548692	550385
ORF	ORF546	ORF547	ORF548	ORF549	ORF550	ORF551	ORF552	ORF553	ORF554	ORF555	ORF556	ORF557	ORF558	ORF559	ORF560	ORF561	ORF562	ORFS63

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-	64	36	45	51	-	_	\downarrow	1	39	88	45	_	39	45	45		_	35	\perp	8
Score	508	353	1324	1009					245	130	519		874	594	334			203		315
Species	Escherichia coli	Haemophilus influenzae	Thermus aquaticus thermophilus	Haemophilus influenzae					Chlamydia psittaci	Chlamydia psittaci	Bacillus subtilis		Escherichia coli	Cucumis sativus	Drosophila melanogaster			Mycobacterium tuberculosis		Chlamydia trachomatis
£	D90832	U32730	U17352	U32824					U65942	U72499	Y13937		U18997	M80571	M58465			294752		M62820
Homology	Tyrosine-specific transport protein (Tyrosine permease).	tyrosine-specific transport protein (tyrP)	L-glutamine:D-fructose-6-P amidotransferase precursor	hypothetical	putative	putative	putative	putative	POMP91A	putative 98 kDa outer membrane protein	putative PlsX protein	putative	ORF f495; orfF of BCMRED, uses 2nd start	glycerol-3-phosphate acyltransferase	insulin-degrading enzyme	putative	putative	unknown	putative	putative heat shock protein ORF; putative
stop	550421	551797	553096	554927	556904	557314	558235	558310	961655	261150	563121	563943	566953	996295	570399	572021	572755	572731	573427	573660
pegin	551611	553041	554946	556300	556524	558126	01873	559215	561349	562931	564083	563593	565379	567079	568021	571269	572519	573519	572879	574160
ORF	ORF564	ORF565	ORF566	ORF567	ORF568	ORF569	ORF570	ORF571	ORF572	ORF573	ORF574	ORP575	ORF576	ORF577	ORF578	ORF579	ORF580	ORF581	ORF582	ORF583

&I	66	39	78	8	37	48	84	66	T		T	T	14	14		42	20	46	42
Score	384	176	358	393	94	695	243	5054					298	339		673	845	719	156
Species	Chlamydia trachomatis	Helicobacter pylori	Chlamydia trachomatis	Chlamydia trachomatis	Bacillus subtilis	Synechocystis sp.	Bacillus subtilis	Chlamydia trachomatis					Staphylococcus aureus	Listeria monocytogenes		Synechocystis sp.	Aquifex pyrophilus	Haemophilus influenzae	Rhodobacter sphaeroides
a	M62820	AE000630	U31570	U31570	X16518	D90899	U87792	U20547					AB001896	U13165		090909	U71154	U32691	U29587
Homology	ribosomal protein S18 homolog; putative	ribosomal protein S6 (rps6)	peptidyl-tRNA hydrolase	peptidyl-tRNA hydrolase	partial ctc gene product (AA 1-186)	glycogen (starch) synthase	phosphatidylglycerophosphate synthase	glycyl-tRNA synthetase	putative	putative	putative	putative	dnaG	DNA primase	putative	DNA mismatch repair protein	DNA mismatch repair protein	excinuclease ABC subunit C (uvrC)	exinuclease ABC subunit C
stop	574184	574446	574923	575057	575469	578023	578017	582104	582206	582811	583182	583438	583827	584299	585016	586610	587758	589408	589578
begin	574426	574781	575243	575458	575849	576545	578673	579012	582697	583122	583514	583869	584435	584967	585297	585240	586484	587786	589198
ORF	ORF584	ORF585	ORF586	ORF587	ORF588	ORF589	ORF590	ORF591	ORF592	ORF593	ORF594	ORF595	ORF596	ORF597	ORF598	ORF599	ORF600	ORF601	ORF602

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L	_	-	ষ্ঠ	_	-	49	23	52	_	88	42	42			_	45	65	37	22	38	1	4
Score			117			991	375	571		1097	242	388				254	638	291	1069	196		780
Species			Bacillus subtilis			Haemophilus influenzae	Synechocystis sp.	Thermus aquaticus thermophilus		Bacillus subtilis	Bacillus subtilis	Escherichia coli				Oryza sativa	Thermoactinomyces intermedius	Arabidopsis thallana	Ricinus communis	Escherichia coli		Helicobacter pylori
A			X62539			U32693	90606G	X70708		X13937	U51868	L14681				U33283	X79068	X57545	L13242	U82598		AE000579
Homology	putative	putative	homologous to E.coli rnpA	putative	putative	cys-tRNA synthetase (cysS)	lysyl-tRNA synthetase	lysinetRNA ligase	putative	putative PriA protein	L-alanine - pimelyl CoA ligase	2-	acylglycerophosphoethanolami ne acyltransferase/acyl carrier protein synthetase	putative	putative	3'(2'), 5-diphosphonucleoside 3'(2') phosphohydrolase	leucine dehydrogenase	inorganic pyrophosphatase	beta-ketoacyl-ACP synthase	HI0034 homolog	putative	conserved hypothetical protein
stop	589630	591272	592765	592849	593121	595637	595640	596154	597282	608009	600734	601910		603531	603757	605610	605582	607493	160809	609296	601019	612927
begin	590061	590739	592406	593145	593900	594138	596122	596864	597731	598524	601876	603523		603794	604413	604549	609909	606843	890609	609652	611860	611812
ORF	ORF603	ORF604	ORF605	ORF606	ORF607	ORF608	ORF609	ORF610	ORF611	ORF612	ORF613	ORF614		ORF615	ORF616	ORF617	ORF618	ORF619	ORF620	ORF621	ORF622	ORF623

81	37	72		47	æ	47	es S	37		4	43	4	45	43
Score	244	134		93	555	448	380	538		313	221	228	543	103
Species	Mycobacterium tuberculosis	Escherichia coli		Mycobacterium tuberculosis	Enterococcus hirae	Haemophilus influenzae	Haemophilus influenzae	Bacillus subtilis		Bacillus subtilis	Escherichia coli	Mycobacterium tuberculosis	Escherichia coli	Bacillus subtilis
A	Z98209	211831		274024	U39788	U32794	U32793	X51419		Y14079	X51584	295388	X51584	AB001488
Homology	trna delta(2) - isopentenylpyrophosphate transferase	delta2- isopentenylpyrophosphate tRNA transferase	putative	unknown	D-alanine:D-alanine ligase	UDP-N-acetylmuramate-alanine ligase (murC)	transferase, peptidoglycan synthesis (murG)	spovE gene product (AA 1-366)	putative	hypothetical protein	murD gene product (AA 1-438)	MurD	ORF-Y (AA 1-360)	PROBABLE UDP-N- ACETYLMURAMOYLALANYL-D- GLUTAMYL-2, 6-DIAMINOLIGASE (EC 6.3.2.15).
stop	612938	613692	615244	615683	615864	617510	618361	619247	620261	620420	621154	621674	622414	623570
begin	613597	613952	614315	615396	617711	618313	619338	620416	619863	621184	621690	622399	623466	624178
ORF	ORF624	ORF625	ORF626	ORF627	ORF628	ORF629	ORF630	ORF631	ORF632	ORF633	ORF634	ORF635	ORF636	ORF637

46	+	8	31				55	47		46	47	47	56		39	46		40	43	42
Score		243	136				467	278		361	460	756	436		1121	1426		416	638	283
Species		Synechocystis sp.	Thermoanaerobacter brockii				Escherichia coli	Escherichia coli		Bacillus subtilis	Bacillus subtilis	Haemophilus influenzae	Rhizobium lequminosarum	1	Thiobacillus ferrooxidans	Haemophilus influenzae		Bacillus subtilis	Salmonella typhimurium	Synechocystis sp.
E		X62437	US6021				U14003	D90837		Z73234	Z73234	U32783	X59956		X95571	U32805		M97208	U06779	D90912
Homology	IIDD-N-acetvimiramovial	glutamyl-2, 6- diaminopimelateD-alanyl-D- alanine ligase	chaperonin 60	putative	putative	putative	elongation factor P	AMP nucleosidase (EC	3.2.2.4).	transketolase	transketolase	transketolase 1 (TK 1) (tktA)	alanyl-tRNA synthetase		alanyl-tRNA synthetase	transcription-repair coupling factor (trcF) (mfd)	putative	uroporphyrinogen decarboxylase	putative oxygen-independent coproporphyrinogen III oxidase	oxygen independent coprophorphyrinogen III
stop	624073		626665	626900	627853	628124	628146	629801		629804	630298	630915	638084		640207	643472	640220	644495	645430	645840
begin	624918		625346	626514	626954	627822	628715	628932		630406	630960	631799	637488		638036	640221	640627	643485	644471	645394
ORF	ORF638		ORF639	ORF640	ORF641	ORF642	ORF643	ORF644		ORF645	ORF646	ORF647	ORF648	00000	OKF649	ORF650	ORF651	ORF652	ORF653	ORF654

1.8 1	88	66	100	100		4	51	52	43	38	36	88	क्ष					\$	51	49	42
Score	133	2043	603	1735		263	456	709	330	405	200	194	205					192	800	288	191
Species	Bacillus subtilis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Haemophilus influenzae	Haemophilus influenzae	Bacillus subtilis	Bacillus subtilis	Bacillus subtilis	Caenorhabditis elegans	Mycobacterium tuberculosis	Helicobacter pylori					Haemophilus influenzae	Arabidopsis thaliana	Saccharomyces cerevisiae	Bacillus subtilis
a	M97208	L25078	M60902	M94254		U32702	U32702	AF008220	M17445	D78193	Z81048	297193	AE000603					U32810	AJ000053	A38767	L09228
Homology	hemY	phosphoprotein	HC1	pCTHoml gene product	putative	phenolhydroxylase component	phenolhydroxylase component	YtpT	spolite protein	Уусл	C41G7.4	hypothetical protein MTCY180.08	D-alanine glycine permease (dagA)	putative	putative	putative	putative	riboflavin synthase beta chain (ribE)	GTP cyclohydrolase II / 3,4-dihydroxy-2-butanone-4-phoshate synthase	unnamed protein product	ribG gene product
stop	647111	647109	650344	651722	652171	652908	653593	661851	662282	663074	663730	663745	664255	665727	665782	668117	668375	668174	668616	669485	866699
begin	645840	649676	649970	650418	651686	652516	62239	659884	661740	662286	662951	664212	665619	666083	666423	666831	668121	668470	669533	669892	670780
ORF	ORF655	ORF656	ORF657	ORF658	ORF659	ORF660	ORF661	ORF662	ORF663	ORF664	ORF665	ORF666	ORF667	ORF668	ORF669	ORF670	ORF671	ORF672	ORF673	ORF674	ORF675

						Transposon Tn5- Mycobacterium tuberculosis Escherichia col Paracoccus denitrificans Bacillus subtil Homo sapiens Helicobacter py	Haloarcula mar. Haloarcula mar. Transposon Th5. Mycobacterium tuberculosis Escherichia col. Paracoccus denitrificans Bacillus subtil Homo sapiens Helicobacter py	Actinopacifius Pleuropneumoniae Haloarcula marisi Transposon Tn5422 Mycobacterium tuberculosis Escherichia coli Paracoccus denitrificans Bacillus subtilis Helicobacter pylo Cyanothece PCC 88 Cyanothece PCC 88	pleuropacifius Pleuropneumoni Haloarcula mar Transposon Tn5 Mycobacterium tuberculosis Sscherichia coi Sacherichia subtil Iomo sapiens Jenitrificans Jenitri	Haloarcula mar. Transposon Th5 Mycobacterium tuberculosis Escherichia col Paracoccus denitrificans Bacillus subtil Homo sapiens Helicobacter py Cyanothece PCC Cyanothece PCC Cyanothece PCC Homo sapiens	leuropheumoni, aloarcula mar aloarcula mar ransposon Th5 ycobacterium uberculosis scherichia col aracoccus arillus subtil pmo sapiens cillus subtil cillus subtil	Haloarcula mari Haloarcula mari Haloarcula mari Transposon Th54 Mycobacterium tuberculosis Escherichia coli Bacillus subtili Homo sapiens Homo sapiens Homo sapiens Homo sapiens Bacillus subtili Bacillus subtili	poscillus poscillus cula mar cula mar cula mar ciclia mar ulosis ichia col ichia col sus subtil bacter py hece PCC hece PCC cus ichia col	non This mar. ila mar. ila mar. ion This osis osis icans icans cter py ce PCC ce PCC ce PCC hia col	sumonida mar. la mar. la mar. srium srium srium srium subtil subtil subtil ia col	eumonida eumonida la mar. la mar. la mar. la mar. la mar. la col lens lens lens lens la col la col la col	eumonida la mar. la mar. la mar. la mar. srium srium subtil subtil subtil ia col
						Halc Tran Mycc tube Esch Bara deni Heli Heli	Halc Tran Mycc tube Esch Baci Homo Heli	Halc Tran Mycc tube Esch Baci Heli Cyan	Halo Tran Tran Sara Sara Sari Saci Saci	Halo Tran Mycc tube Esch Baci Homo Heli	allo and allo allo allo allo allo allo allo all			Narcullius sap	Ispose ispose ispose ispose ispose ispose ispose ispose is interior interior interior is interior i	Ispose is	nspose ispose is
						Myc tub Esc Esc den den Hen Hel	Tra Myc tub Esc Esc den Hom Hel	Tra Myc tub Ryc Esc Bsc Hom Hell	Yaa len len lac	Myc tub Bac			oba	nspos obact ercul heric itilius o sap icoba icoba illus o sap	nsposc obacte obacte herich itrifi illus copac icobac	nspose obacte ercule hericl illus illus cobac nothec	nspose obacte erculc hericl itrifi illus cobacc nothec nothec
157	17 15											Tran Myco tube Esch Baci Heli Baci Homo	Transp Mycoba tuberc Escher Paraco denitr Bacill Homo s Homo s	Transpos Mycobact tubercul Escheric Paracocc denitrif Bacillus Helicoba Helicoba Homo sap Homo sap	Transpose Mycobacte tuberculc Escherich Baracoccu denitrifi Bacillus Homo sapi Homo sapi	Transpose Mycobacte tubercule Eschericl Paracocce denitrifi Bacillus Homo sapi Homo sapi Homo sapi	Transpose Mycobacte tuberculc Escherich Paracoccu denitrifi Bacillus Homo sapi Homo sapi
E13 18 18	310 402 285 235	3104 1025 2857 2354 9228	1025 1025 2857 2354 2358 3068	L28104 Z74025 M22857 L02354 L09228 M23068	3104 1025 2857 2354 3068 000542	3104 1025 2857 2354 3228 3068 000542	3104 1025 2857 2354 3068 3068 300542	2857 2857 2354 2358 3068 3068 3068 3068 3068 3068	80 0	80 0	880	B 8 0 0 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	80 87	880	8 8 8 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	B 8 4 5 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	B 8 4 5 5 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7
Z740 M228 L023	Z7402 M2285 L0235	Z74025 M22857 L02354 L09228	Z74025 M22857 L02354 L09228 M23068	Z74025 M22857 L02354 L09228 M23068 AE000542	M22857 L02354 L09228 M23068 AE000542	M22857 L02354 L09228 M23068 AE00054	Z74025 M22857 L02354 L09228 M23068 AE00054	Z74025 M22857 L02354 L09228 M23068 AE00054 AF00178	Z74025 M22857 L02354 L09228 M23068 AE00054 AF00178	AF00178 AF00178 AF00178 A72718	Z74025 M22857 L02354 L09228 M23068 AE00054 AF00178 M72718	AE00059 M72857 L02354 L09228 M23068 AE00054 AF00178 M72718 M72718	Z74025 M22857 L02354 L09228 M23068 AE00054 AF00178 S73498	Z74025 M22857 L02354 L09228 M23068 AE00054 AF00178 S73498 U00039	AF00178 AF00178 AF00178 A72718 TO0039	AF00178 AF00178 AF00178 A72718 TO0039	Z74025 M22857 L02354 L09228 M23068 AE00178 AF00178 S73498
unknown rod-shape-determining protein biotin [acetyl-CoA	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase	unknown rod-shape-determining protein biotin [acety]-CoA carboxylase] ligase ORFX13	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs)	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis,	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa]	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) Nifu putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, infertile, 505 aa] L-glycerol 3-phosphate dehydrogenase	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, heptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative putative putative putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) Nifu putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative putative putative putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Feptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative putative putative putative putative putative	unknown rod-shape-determining protein biotin [acetyl-CoA carboxylase] ligase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Feptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative
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676463 rod-s prote 676476 bioti	·																
			+++	676476 677700 679508 680502	676476 677700 679508 680502 681280	676476 677700 679508 680502 681280 682558						+++++++++++++++++++++++++++++++++++++++					
biotin [acety	biotin [acety carboxylase]																
	carboxylase] 11gase	Carboxylase Ingase ORFX13	ORFX13 2,3-bisphosphoglycerate	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs)	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifS)	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative	Carboxylase] iigase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifS) NifU putative putative	Carboxytasel ingase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4	Carboxytasel iigase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifS) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis,	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Feptide, 505 aa]	ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase	Carboxytasel iigase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative	Carboxytasel ilgase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifS) NifU putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Feptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative	Carboxylasel ilgase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifS) NifU putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative putative putative	Carboxylasel ilgase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) NifU putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative putative putative putative putative putative	Carboxylasel ilgase ORFX13 2,3-bisphosphoglycerate synthesis of [Fe-S] cluster (nifs) Nifu putative ORF 4 putative AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] L-glycerol 3-phosphate dehydrogenase putative

8	202	ŝ	26	50			न्न	23	55	88	41		4	42	28	69	42		66	22	45					
Score	1010	0101	196	1073			84	615	1183	362	809		165	155	1044	258	179		1548	713	273		-			
Species	Neoral limetiv	frontalis	Bacillus subtilis	Bacillus cereus			Helicobacter pylori	Myxococcus xanthus	Escherichia coli	Bacillus subtilis	Synechocystis sp.		Cyanidium caldarium	Bacillus firmus	Mycobacterium smegmatis	Zea mays	Streptomyces lividans		Chlamydia trachomatis	Bacillus subtilis	Micrococcus luteus					
A	M59372		M96343	X98455			AE000591	AF013216	L18867	L47709	D90912		AF022186	X99401	U66081	U71123	U21192		U72715	X62539	U22181					
Homology	phosphoenolpyruvate	carboxykinase	MreB protein	SNF	putative	putative	trigger factor (tig)	proteosome major subunit	ATP-dependent protease ATPase subunit	poly(A) polymerase	hypothetical protein	putative	Preprotein translocase subunit	весд	SecA	cp-SecA; chloroplast SecA homolog	SecA	putative	phosphatidylserine decarboxylase	homologous to E.coli 50K	ultraviolet N-glycosylase/AP lyase	putative	putative	putative	putative	putative
Btop	692041		693837	694942	021169	626269	700117	700895	702165	703412	705000	705604	705704	706138	706496	708078	708248	708872	710262	712763	713438	713651	714120	714834	715558	715921
pegin	693837		694934	697263	698084	698392	698792	700269	700912	702183	703522	705011	706159	706521	708103	708398	708610	710278	711164	711432	712767	714232	714632	715592	715854	716937
ORF	ORF700		ORF701	ORF702	ORF703	ORF704	ORF705	ORF706	ORF707	ORF708	ORF709	ORF710	ORF711	ORF712	ORF713	ORF714	ORF715	ORF716	ORF717	ORF718	ORF719	ORF720	ORF721	ORF722	ORF723	ORF724

# #	92		42		37	1	4	25	39	14	22	88	71	83		33	43	66	92	06	8
Score	2049		766	-	164	722	477	388	805	+	607	366	854 7	531		115 3	208 4	2045 9	1269 7	1278 9	1153 9
Sc	2	+-	 	-		7,	4	ļ	 	17)9	26	8	53	-	11	ļ	70	12	12	17
Species	Chlamydia trachomatis		Haemophilus influenzae		Haemophilus influenzae	Bacillus subtilis	Escherichia coli	Caulobacter crescentus	Escherichia coli	Synechocystis sp.	Escherichia coli	Synechocystis sp.	Synechocystis sp.	Synechocystis sp.		Saccharomyces cerevisiae	Haemophilus influenzae	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis
A	U83197		U32834		U32834	X56678	U00039	U87804	D90811	D64004	D90811	D64004	D64004	D64004		L36940	U32688	M14738	U60196	U60196	U60196
Homology	3-phosphoglycerate kinase	putative	phosphate permease (YBR296C)	putative	H. influenzae predicted coding region HI1603	dciAD	was dppE	chromosome partitioning protein ParB	NifS protein.	hypothetical protein	Multidrug resistance protein 1 (P-glycoprotein 1).	ABC transporter subunit	ABC transporter subunit	ABC transporter subunit	putative	antiviral protein	penicillin-binding protein 2 (pbp2)	major outer membrane protein precursor	ribosomal protein S2	elongation factor Ts	UMP kinase
stop	717149	718862	718499	719782	720144	721575	722356	722397	723378	724576	725767	726538	726753	727469	728329	728759	729442	734427	735659	736504	737254
pegin	718357	718500	719797	720273	720452	720613	721559	723248	724598	725763	726519	726819	727493	727984	728778	729346	732639	733246	734814	735644	736520
ORF	ORF725	ORF726	ORF727	ORF728	ORF729	ORF730	ORF731	ORF732	ORF733	ORF734	ORF735	ORF736	ORF737	ORF738	ORF739	ORF740	ORF741	ORF742	ORF743	ORF744	ORF745

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å.	92	1	3	37		1	8	88		\perp			55	\perp						40	æ	49	8
Score	760		116	453			137	117					1220							172	970	409	107
Species	Chlamydia trachomatis		Listeria monocytogenes	Pseudomonas aeruginosa			Mycobacterium tuberculosis	Caulobacter crescentus		·			Pseudomonas aeruginosa							Bacillus subtilis	Clostridium acetobutylicum	Mycobacterium smegmatis	Mycobacterium tuberculosis
Ω	96T09N		U40604	X68594			Z80233	M69228					AF010151							M57676	U35453	X92503	274024
Homology	ribosome-releasing factor	putative	ORF3; putative 39 kDa protein	ХсрQ	putative	putative	unknown	putative	putative	putative	putative	putative	PscN	putative	putative	putative	putative	putative	putative	NAD(P)H:glutamyl-transfer RNA reductase	DNA gyrase subunit B	gyrA	unknown
stop	737787	738679	739740	740060	742045	742824	743306	744430	744611	744958	745561	746416	746944	748274	748965	749433	749721	750007	752503	753616	756814	758301	758446
begin	737254	737942	738838	742057	742869	743378	744298	744714	744985	745557	746412	746772	748269	748966	749426	749702	750029	752307	752913	754659	755000	756796	758691
ORF	ORF746	ORF747	ORF748	ORF749	ORF750	ORF751	ORF752	ORF753	ORF754	ORF755	ORF756	ORF757	ORF758	ORF759	ORF760	ORF761	ORF762	ORF763	ORF764	ORF765	ORF766	ORF767	ORF768

	7	T	т-			1		_	,														
*	48			8	8	66	5		66	1 00	100	66	100					35	51	52			40
Score	241			1350	536	1197	239		1803	704	1753	904	2249					486	263	1357			93
Species	Escherichia coli			Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis					Bacillus subtilis	Escherichia coli	Bacillus subtilis			Escherichia coli				
£	U50134			U72493	U72493	U72493	M17875		232530	Z32530	U16739	Z32530	232530					D26185	U18997	L27797			U29581
Homology	SfhB	putative	putative	3-deoxy-D-manno-octulosonate 8-phosphate synthetase	unknown	ATP binding protein	chlanectin coding region	putative	unknown function	unknown function	RecA	unknown function	unknown function	putative	putative	putative	putative	unknown	ORF_f169	DNA topoisomerase I	putative	putative	ORF_f397
stop	759338	759871	760188	761772	762142	762983	763335	764438	764821	766065	766934	768252	768791	770470	770185	770634	771330	773391	773427	774191	777706	776953	777732
begin	759787	760242	760538	996094	761759	762267	764465	764857	766068	766643	768091	768785	770092	770138	770661	770924	772010	772390	774221	776035	776663	777195	779222
ORF	ORF769	ORF770	ORF771	ORF772	ORF773	ORF774	ORF775	ORF776	ORF777	ORF778	ORF779	ORF780	ORF781	ORF782	ORF783	ORF784	ORF785	ORF786	ORF787	ORF788	ORF789	ORF790	ORF791

Homology ID
putative
exonuclease V (AA 1-1180) X04581
putative
MreC protein
aspartate aminotransferase M12105 precursor
putative
U02878
putative
NADH:ubiquinone
Oxidoreductase subunit A
dehydratase
putative
putative
putative
ompR gene product
glucose-1-phosphate
thymidylyltransferase
farnesyl diphosphate
synthase
putative
Orf39.9
This ORF is homologous to a
40.0 kd hypothetical protein
in the htrB 3' region from
E. coli, Accession Number

1.8 1.8	46	32	97	97	9	29	Τ	SS	Π	Γ	Ī		Γ						5	~	9				<u></u>
-			ļ	"		2	-	2	\vdash	_	\vdash	\vdash	\vdash	-	-	_	-	L	88	32	=	_	45		48
Score	407	397	1716	973	280	775		125											150	90	2868		310	•	100
Species	Helicobacter pylori	Caenorhabditis elegans	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia pneumoniae		Chlamydia pneumoniae											Odontella sinensis	Homo sapiens	Chlamydia trachomatis		Pisum sativum		Neisseria meningitidis
αī	AE000633	U40707	U50732	U50732	U50732	L23921		L23921											Z67753	AB002334	J05546		X08611		X68068
Homology	ribosomal protein S4 (rps4)	apurinic/apyrimidinic endonuclease	mviB homolog	mviB homolog	lorf2; possible membrane- bound protein	76 kDa protein	putative	76 kDa protein	putative	30S ribosomal protein S20	KIAA0336	RNA polymerase sigma-subunit	putative	dihydropterin	pyrophosphokinase /dihydropteroate synthase	dihydropteroate synthase									
stop	799927	800029	802129	802673	803246	804220	805356	806282	808081	809009	809079	810328	812342	813522	813772	814334	814314	814396	815428	817456	819320	819713	820402		821061
begin	799301	800892	801062	802023	802851	803105	804307	805290	806453	808026	810461	811605	811725	812329	813455	813732	815213	814878	815733	816116	817608	819324	819704		820375
ORF	ORF813	ORF814	ORF815	ORF816	ORF817	ORF818	ORF819	ORF820	ORF821	ORF822	ORF823	ORF824	ORF825	ORF826	ORF827	ORF828	ORF829	ORF830	ORF831	ORF832	ORF833	ORF834	ORF835		ORF836

	Increase	
ase Z84379 Streptococcus	ıctas	dihydrofolate reductase
ted U67522 Methanococcus	predicted MJ0768	M. jannaschii predic coding region MJ0768
		putative
M58480 Thiobacillus ferrooxidans	 =	nitrogen metabolism regulator
M63176 Staphylococcus		helicase
M63176 Staphylococcus		helicase
X73124 Bacillus subtilis	נו	ipa-57d gene product
		putative
U32712 Haemophilus influenzae		hypothetical
stretch (32- L19954 Bacillus subtilis to N-terminal sequence of y cloned B.	ch terr uenc ned ativ	19/20 residue stretch (32 51) identical to N-termin putative signal sequence unknown, partly cloned B. subtilis gene.; putative
	iroE	heat shock protein GroEL
. Z34917 Hordeum vulgare		bas1 protein
		putative

ORF	pegin	stop	Homology	Ð	Species	Score	*
ORF858	837209	838699	putative				
ORF859	838760	839575	putative				
ORF860	839942	840583	putative				
ORF861	840445	841713	putative				
ORF862	841659	842459	putative				
ORF863	842523	843068	putative				
ORF864	843495	843031	putative				
ORF865	843239	846196	putative				
ORF866	844137	843802	putative				
ORF867	848043	846217	putative				
ORF868	850123	848150	putative				
ORF869	851645	850230	putative				
ORF870	853696	851669	putative				
ORF871	854836	853700	putative				
ORF872	855525	854920	putative				
ORF873	856240	855437	putative				
ORF874	857183	856233	putative				
ORF875	859439	857451	putative				
ORF876	859946	859587	putative				
ORF877	859642	860640	putative				
ORF878	861599	860724	putative				
ORF879	862053	861580	putative				
ORF880	863540	862098	putative				Ī
ORF881	863930	863571	putative				
ORF882	864697	966898	putative				
ORF883	864923	866248	DNA mismatch repair protein (mutL)	U32692	Haemophilus influenzae	206	47
ORF884	866303	866605	putative				
ORF885	866665	867732	YqhT	D84432	Bacillus subtilis	444	39
ORF886	867810	060698	putative			-	
ORF887	869094	869357	putative				
ORF888	869270	871372	fimbrial assembly protein	L13865	Pseudomonas aeruginosa	181	40

8 T 8	999	84	33					8	4	46		8		43	4	æ	4		43	35	41		_
Score	825	213	271					174	172	464		141		236	685	578	437		824	230	220		
Species	Xanthomonas campestris	Acinetobacter calcoaceticus	Escherichia coli					Salmonella typhimurium	Yersinia pseudotuberculosis	Xanthomonas campestris		Pseudomonas aeruginosa		Erwinia amylovora	Saccharomyces cerevisiae	Clostridium magnum	Bacillus subtilis		Synechocystis sp.	Clostridium perfringens	Bacillus subtilis		
Ð	X59079	Y09102	U18997					X99944	125667	M64094		U56077		U56662	275104	L31844	D84432		D90916	D49784	X13937		
Homology	xpsE gene product	secretion protein XcpR	ORF_0398	putative	putative	putative	putative	secretion system apparatus, SsaT	yscS	pathogenicity protein	putative	PscL	putative	HrcJ	ORF YOR196C	dihydrolipoamide dehydrogenase	Λįτλ	putative	helicase of the snf2/rad54 family	sodium-coupled branched- chain amino acid carrier	putative Fmu protein	putative	
stop	872582	872860	873915	873360	874438	875386	876382	877000	877876	878172	879161	879105	880052	88088	881948	882901	883661	884508	885166	888940	890325	891116	
begin	871299	872429	872773	873812	874028	874778	875774	877872	878172	843628	878883	879842	880885	881863	882904	883794	884296	884996	888777	890172	891164	891463	
ORF	ORF889	ORF890	ORF891	ORF892	ORF893	ORF894	ORF895	ORF896	ORF897	ORF898	ORF899	ORF900	ORF901	ORF902	ORF903	ORF904	ORF905	ORF906	ORF907	ORF908	ORF909	ORF910	

18			39				43		52	47			51	86	100	32	9	ļ	37	40		53		S		ස	59
Score			155				1974		1117	989			1339	1196	607	000	360		150	181		197		145		309	302
Species			Synechocystis sp.				Helicobacter pylori	1	Escherichia coli	Streptococcus	equisimilis		Synechocystis sp.	Chlamydia trachomatis	Chlamydia trachomatis	Chlamidia neittani			Synechocystis sp.	Staphylococcus aureus		Helicobacter pylori		Sinorhizobium meliloti		Pseudomonas aeruginosa	Synechocystis sp.
αI			D90908				AE000646		M17102	217214			D90910	X66126	140369	1.39892	70000	, 10000	D90914	Z48003		U94318		L13845		U67855	D90910
Homology	putative	putative	hypothetical protein	putative	putative	putative	DNA polymerase III alpha-	subunit (dnaE)	UhpC protein	histidinetRNA ligase		putative	aspartyl-tRNA synthetase	mip-like protein	nods	trxA	nitative	hymothetical arcteia	nyponierical procein	DNA polymerase III	putative	VdlD	putative	acid-inducible gene	putative	UDP-3-0-acyl-GlcNAc deacetylase	(3R) -hydroxymyristol acyl carrier protein dehydrase
stop	893808	893643	893821	894248	895050	896829	897064		900791	903876		903471	905605	906474	906945	907001	908742	909194	*CTC0C	909584	909951	910569	910944	912261	912629	913218	913676
begin	893356	893909	894276	894778	895892	895951	900783		902032	902659		903731	903860	905725	906493	907306	101806	908721	17/000	901006	909583	910081	910615	910948	912399	912595	913203
ORF	ORF912	ORF913	ORF914	ORF915	ORF916	ORF917	ORF918		ORF919	ORF920		ORF921	ORF922	ORF923	ORF924	ORF925	ORF926	ORP927	12CINO	ORF928	ORF929	ORF930	ORF931	ORF932	ORF933	ORF934	ORF935

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*H	8	45		_	_	84	43	47	48	8	49	25	62	66	<u>5</u>	9	100	66
Score	503	407				470	210	116	800	315	240	909	434	343	419	618	268	793
Species	Rickettsia rickettsii	Escherichia coli				Haemophilus influenzae	Synechococcus sp.	Thermotoga maritima	Mycoplasma-like organism	Mycoplasma pneumoniae	Thermotoga maritima	Haemophilus influenzae	Thermotoga maritima	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis
αī	רב2690	X63666				U32761	AB000111	221677	M74770	AE000061	Z21677	U32761	221677	M80325	M80325	M80325	M80325	M80325
Homology	UDP-N-acetylglucosamine acyltransferase	methionyl-tRNA formyltransferase	putative	putative	putative	ribosomal protein L3 (rpL3)	50S ribosomal protein L4	ribosomal protein L23	rpl2	Mycoplasma pneumoniae, ribosomal protein S19; similar to GenBank Accession Number S36895, from M. bovis	ribosomal protein L22		ribosomal protein L16	ribosomal protein CtrL29e	ribosomal protein S17e	ribosomal protein CtrL14e	ribosomal protein CtrL24e	ribosomal protein CtrL5e
stop	914485	915136	915467	916633	916539	917627	918304	918655	919533	919829	920157	920840	921294	921514	921758	922143	922491	923035
begin	913691	914516	915144	915629	916051	916965	917612	918323	918682	919542	919723	920184	920866	921272	921510	921778	922159	922496
ORF	ORF936	ORF937	ORF938	ORF939	ORF940	ORF941	ORF942	ORF943	ORF944	ORF945	ORF946	ORF947	ORF948	ORF949	ORF950	ORF951	ORF952	ORF953

% H	86	8	66	86	66	66	100	86	97	86	66	Τ	Τ			14	38	09
Score	487	927	605	814	740	2254	604	646	847	1040	1735					250	258 .	264
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis					Haemophilus influenzae	Haemophilus influenzae	Helicobacter pylori
a	M80325	M60652	M80325	M80325	M80325	L25077	L33834	L33834	L33834	L33834	U83198					U32717	U32716	AE000540
Homology	ribosomal protein CtrS8e	ribosomal protein L6	ribosomal protein CtrL18e	ribosomal protein CtrS5e	ribosomal protein CtrL15e	homolog	ribosomal protein S13	ribosomal protein S11	RNA polymerase alpha-subunit	RNA polymerase alpha-subunit	glyceraldehyde-3-phosphate dehydrogenase	putative	putative	putative	putative	crossover junction endodeoxyribonuclease (ruvC)	Holliday junction DNA helicase (ruvA)	nucleoside diphosphate kinase (ndk)
stop	923453	924032	924425	924937	925364	926760	927184	927604	928155	928759	930244	930656	931078	931666	931959	932579	933201	933621
pegin	923160	923484	924048	924443	924933	925390	926819	927209	927577	928100	929222	930222	809086	931367	931549	932070	932602	933319
ORF	ORF954	ORF955	ORF956	ORF957	ORF958	ORF959	ORF960	ORF961	ORF962	ORF963	ORF964	ORF965	ORF966	ORF967	ORF968	ORF969	ORF970	ORF971

1%	2		36	51	41	33	40	36	42	32	8	89	4		41		42	47			T
Score	186		156	1562	848	120	899	265	1334	198	882	417	755		223		260	178			
Species	Myxococcus xanthus		Mycoplasma genitalium	Pseudomonas putida		Helicobacter pylori	Arabidopsis thaliana	Bacillus subtilis	Bacillus caldotenax	Caenorhabditis elegans	Pseudomonas fluorescens	Deinococcus radiodurans	Oryza sativa		Bacillus subtilis		Mus musculus	Cucumis sativus			
ΩI	J05207		U39706	X62540	D26185	AE000610	Z49227	AF008220	D12982	227079	L27278	L27276	M31616		298682		M59288	D26106			
Homology	nucleoside 5'-diphosphate phosphotransferase (EC	2.7.4.6)	hypothetical protein (GB:U14003_297)	homologous to E.coli gidA	replicative DNA helicase	phosphatidylglycerophosphate synthase (pgsA)	adenine nucleotide translocase	putative protease	DNA polymerase	T05G5.5	The first ATG in the open reading frame was chosen as the initiation codon.	'The first GTG in the open reading frame was chosen as the initiation codon.'	ADPglucose pyrophosphorylase	putative	YlbH protein	putative	ferrochelatase	ferrochelatase	putative	putative	putative
stop	933785		933848	934539	999986	860686	940933	942068	944685	945287	946294	946676	948454	949277	949594	950676	951330	951643	952798	954264	955157
begin	933522		934546	936377	938081	938538	939329	941031	942082	944634	945287	946293	947105	948522	949277	949849	950680	951281	951788	953581	954426
ORF	ORF972		ORF973	ORF974	ORF975	ORF976	ORF977	ORF978	ORF979	ORF980	ORF981	ORF982	ORF983	ORF984	ORF985	ORF986	ORF987	ORF988	ORF989	ORF990	ORF991

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1%	41	38	98 98	39			37		4			45		30		20	44	42	39		4	52	T
Score	130	327	263	109			775		492			849		147		500	141	142	183	-	534	187	
Species	Methanosarcina barkeri	Bacillus subtilis	Helicobacter pylori	Methanococcus jannaschii			Bacillus subtilis		Saccharomyces cerevisiae			Pseudomonas aeruginosa		Myxococcus xanthus		Bacillus subtilis	Bacillus subtilis	Bscherichia coli	Saccharomyces cerevisiae		Bacillus firmus	Bacillus subtilis	
Ð	X93084	X56347	AE000548	U67593			275208		248008			212154		L39904		D64126	D26185	AE000126	Z49939		X99401	AF013188	
Homology	orf4 gene product	OppB gene product	dipeptide ABC transporter, permease protein (dppC)	methylated DNA protein cysteine methyltransferase	putative	putative	phenylalanyl-tRNA synthetase beta subunit	putative	unknown	putative	putative	transcriptional activator of	pilA	sensor protein	putative	unknown	unknown	hypothetical protein in htrA dapD intergenic region	unknown	putative	peptide release factor 2	release factor 2	putative
stop	957940	959312	961050	961053	961487	961584	962545	965708	966193	968061	968064	969528		971024	972388	973746	974558	975207	976254	976899	977635	977933	978433
begin	955754	957837	959299	961562	962575	961949	964914	964941	967023	967444	968903	970685		971806	973053	974546	975223	976193	976520	976588	976886	977661	977918
ORF	ORF992	ORF993	ORF994	ORF995	ORF996	ORF997	ORF998	ORF999	ORF1000	ORFIO01	ORF1002	ORF1003		ORF1004	ORF1005	ORF1006	ORF1007	ORF1008	ORF1009	ORF1010	ORF1011	ORF1012	ORF1013

1%	52	9				88		4	88	35	SS SS	46			69		æ		38		£3			98	75	83
Score	355	199				593		347	782	224	286	132			 343		110		447		240		+	880 ,	533	2018
Species	Bacillus subtilis	Haemophilus influenzae				Haemophilus influenzae		Synechocystis sp.	1	Synechocystis sp.	Mycobacterium leprae	Escherichia coli			Helicobacter pylori	1	Methanococcus jannaschii		Synechocystis sp.		Rhodobacter sphaeroides			Chlamydia psittaci	Chlamydia psittaci	Chlamydia psittaci
a	D50551	U32717				U32788		D64006	D64006	D90915	U00021	AE000238			AE000627		U67577		D64000		U83136			U41759	U41759	U41759
Homology	spore coat protein CotRC	hypothetical	putative	putative	putative	UDP-N-acetylglucosamine	<pre>enolpyruvyl transferase (murZ)</pre>	arginyl-tRNA-synthetase	arginyl-tRNA-synthetase	hypothetical protein	No definition line found		pct identical (24 gaps) to	256 aa protein CDSA RCOLT	conserved hypothetical	protein	hypothetical protein (HI0920)	putative		process section	protein-export membrane protein	putative	putative	RecJ recombination protein	unknown	glutamyl-tRNA synthetase homolog
stop	978984	979331	979389	980112	981148	983598		983862	984371	985399	986046	986693			987616		987936	989163	993442	10000	993785	993416	994262	995656	119966	998267
pegin	978619	978933	261186	111616	982116	982321		984488	985381	986103	986693	987607			988119		988253	988831	989693	007.00	8042	993835	993882	994226	98036	996885
ORF	ORF1014	ORF1015	ORF1016	ORF1017	ORF1018	ORF1019		ORF1020	ORF1021	ORF1022	ORF1023	ORF1024			ORF1025		ORF1026	ORF1027	ORF1028	9001990	OKF 1029	ORF1030	ORF1031	ORF1032	ORF1033	ORF1034

% H	100	100	\$	97	42	02	8	83	88	48	42	22		4		6		
Score	504	2857	276	438	486	454	662	2147	350	113	102	1021		365		108	-	
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Escherichia coli	Haemophilus influenzae	Chlamydia trachomatis	Helicobacter pylori	Spirulina platensis	Bscherichia coli	Mycobacterium tuberculosis	Bacillus subtilis		Synechocystis sp.		Haemophilus influenzae		
CI CI	M35148	M23001	M35148	M35148	D00674	M86701	211567	AE000625	221676	M23008	Z92774	238002		D90915		U32759		
Homology	9-kDa cysteine-rich outer membrane protein	outer membrane protein 2	15-kDa serine-rich outer membrane protein	15-kDa serine-rich outer membrane protein	ORF of prc gene (alt.)	StrA	ribosomal protein S7	translation elongation factor EF-G (fusA)	ribosomal protein S10	NADPH-sulfite reducatase flavoprotein component	unknown	serine hydroxymethyltransferase	putative	ATP-dependent Clp protease proteolytic subunit	putative	diaminopimelate epimerase (dapF)	putative	putative
stop	999225	1001033	1001516	1001664	1001823	1004845	1005382	1007496	1007821	1008698	1009121	1012054	1011942	1012635	1012862	1013440	1014055	1014489
pegin	296866	999375	1001211	1001392	1003721	1004459	1004990	1005391	1007486	1007802	1009426	1010534	1012397	1012042	1012593	1012811	1013456	1013977
ORF	ORF1035	ORF1036	ORF1037	ORF1038	ORF1039	ORF1040	ORF1041	ORF1042	ORF1043	ORF1044	ORF1045	ORF1046	ORF1047	ORF1048	ORF1049	ORF1050	ORF1051	ORF1052

%I	38			42					33	36				43		2	51	37	35	4	35	49		36			46	
Score	263			428					164	201				218		251	603	161	439	312	354	95		75			160	
Species	Bscherichia coli			Helicobacter pylori					Helicobacter pylori	Mycobacterium	tuberculosis			Escherichia coli		Porphyra purpurea	Escherichia coli	Haemophilus influenzae	Synechocystis sp.	Bacillus subtilis	Synechocystis sp.	Chlamydia psittaci		Chlamydia psittaci			Chlamydia psittaci	
Ð	AE000459			AE000579					AE000647	295208				U18997		U38804	U18997	U32769	D90903	AB000617	D90903	U72499		U72499			U72499	
Homology	hypothetical 28.1 kD protein in udp-rfaH intergenic	region	putative	conserved hypothetical	protein	putative	putative	putative	hemolysin	unknown		putative	putative	50S ribosomal subunit	protein L21	50S ribosomal protein L27	ORF_f390	GTP-binding protein (obg)	hypothetical protein	YedI	adhesion protein	putative 98 kDa outer	membrane protein	putative 98 kDa outer	membrane protein	putative	putative 98 kDa outer	putative
stop	1014529		1015145	1015939		1017245	1017916	1018580	1019831	1020114		1021075	1022097	1023667		1023949	1024776	1025045	1024967	1025839	1026546	1027929		1030508		1032086	1033456	1035910
begin	1015224		1016002	1017120		1017766	1018911	1019191	1020199	1021007		1021569	1022411	1023344		1023701	1023976	1024704	1025881	1026546	1027379	1030604		1033252		1031733	1037037	1035674
ORF	ORF1053		ORF1054	ORF1055		ORF1056	ORF1057	ORF1058	ORF1059	ORF1060		ORF1061	ORF1062	ORF1063		ORF1064	ORF1065	ORF1066	ORF1067	ORF1068	ORF1069	ORF1070		ORF1071		ORF1072	ORF1073	ORF1074

Score I%	
Species	
a	
Homology	putative
gctep	1036507
pegin	1036175
ORF	ORF1075

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¥ H		97		95	94	100	100		86	66	100	86			94			97			
Score		658		284	303	501	476		583	607	534	529			395			631	-		
Species		Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis			Chlamydia trachomatis			Chlamydia trachomatis			
Ωī		AE001313		AE001309	AE001309	AE001309	AE001308		AE001307	AE001307	AE001307	AE001305			AE001305			AE001303			
Homology	putative	<pre>GutQ/KpsF Family Sugar-P Isomerase</pre>	putative	hypothetical protein	hypothetical protein	hypothetical protein (possible 357R?)	Predicted OMP [leader (19) peptide]	putative	hypothetical protein	Lon ATP-dependent protease	hypothetical protein	hypothetical protein	putative	putative	hypothetical protein	putative	putative	predied ferredoxin	putative	putative	putative
stop	1036967	16989	31408	56834	56913	65074	73195	78736	79411	81959	86999	109456	111351	111734	112911	113715	120464	125522	131604	144427	150369
pegin	(woo) 89	16291	31779	56502	26686	64748	73482	78482	79803	82333	87313	109929	111599	112069	112666	114017	120757	125133	131888	144164	150698
ORF	ORF1076	ORF1077	ORF1078	ORF1079	ORF1080	ORF1081	ORF1082	ORF1083	ORF1084	ORF1085	ORF1086	ORF1087	ORF1088	ORF1089	ORF1090	ORF1091	ORF1092	ORF1093	ORF1094	ORF1095	ORF1096

124 124 124 124 124 124 124 124 124 126	ORF	begin	stop	Homology	QI	Species	Score	4.4
168742 168154 hypothetical protein AE001300 Chlamydia trachomatis 724 168742 168425 hypothetical protein AE001300 Chlamydia trachomatis 356 1701509 170793 hypothetical protein AE001209 Chlamydia trachomatis 489 177145 177474 Carboxylaee/Transferase AE001299 Chlamydia trachomatis 518 188295 188023 hypothetical protein AE001298 Chlamydia trachomatis 733 188791 188330 hypothetical protein AE001298 Chlamydia trachomatis 733 190629 190336 putative AE001299 Chlamydia trachomatis 733 197313 197083 putative AE001299 Chlamydia trachomatis 507 235160 234852 Glutamate Aminomutase AE001295 Chlamydia trachomatis 507 249733 249446 Oligopeptide Permease AE001293 Chlamydia trachomatis 1860 253701 254789 hypothetical protein AE001293 Chlamydia trachomatis 1860 271633 271932 hypothetical protein AE001291 Chlamydia trachomatis 700 527566 276070 Oxidoreductase AE001291 Chlamydia trachomatis 700 527581 2778218 putative AE001291 Chlamydia trachomatis 700 527591 2778218 putative AE001291 Chlamydia trachomatis 700 527791 2778218 putative AE001291 Chlamydia trachomatis 700 527791 2778218 putative AE001291 Chlamydia trachomatis 700 527791 2778218 putative AE001291 Chlamydia trachomatis 700 57791 700 57791 700 7	ORF1097	164385	163948	NADH (Ubiquinone) Dehydrogenase	AE001300		755	100
168742 168425 hypothetical protein AB001300 Chlamydia trachomatis 356 170569 170793 Accordant Ab001300 Chlamydia trachomatis 518 177145 Accordant Ab001299 Chlamydia trachomatis 518 188295 188023 hypothetical protein AB001299 Chlamydia trachomatis 518 188791 188330 hypothetical protein AB001299 Chlamydia trachomatis 733 188792 19083 putative Ab001299 Chlamydia trachomatis 733 190629 190336 putative Ab001299 Chlamydia trachomatis 507 230720 23493 Dutative AB001295 Chlamydia trachomatis 507 249733 249446 Oligopetide Permease AB001293 Chlamydia trachomatis 1860 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 1860 271633 271932 Disulfide bond AB001291 Chlamydia trachomatis 700 517931 27566 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700 517931 277567 275216 Dutative AB001291 Chlamydia trachomatis 700 517931 277567 275266 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700 517931 277531 278216 Dutative AB001291 Chlamydia trachomatis 700 517931 277561 278216 Dutative AB001291 Chlamydia trachomatis 700 517931 277511 278216 Dutative AB001291 Chlamydia trachomatis 700 700 277511 278216 Dutative AB001291 Chlamydia trachomatis 700 700 277511 278216 AB001291 AB00129	ORF1098	165690	166115		AE001300		724	66
170569 170793 hypothetical protein AB001299 Chlamydia trachomatis 489 177145 177474 Carboxylase/Transferase AB001299 Chlamydia trachomatis 518 188295 188023 hypothetical protein AB001298 Chlamydia trachomatis 733 188791 188330 hypothetical protein AB001298 Chlamydia trachomatis 733 190629 190336 putative AB001299 Chlamydia trachomatis 507 237127 236913 putative AB001295 Chlamydia trachomatis 507 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 7 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 1860 5 271633 271932 hypothetical protein AB001291 Chlamydia trachomatis 512 7 275566 275070 Oxidoreductase AB001291 Chlamydia trachomatis 700 5 277931 278218 putative Dieulfide bond AB001291 Chlamydia trachomatis 700 5 277931 278218 putative Dieulfide bond AB001291 Chlamydia trachomatis 700 5 277931 278218 putative Dieulfide bond AB001291 Chlamydia trachomatis 700 5 277931 278218 putative Dieulfide bond AB001291 Chlamydia trachomatis 700 5 277931 278218 putative Dieulfide bond AB001291 Chlamydia trachomatis 700 5	ORF1099	168742	168425		AE001300	1	356	96
177145 177474 CacoA AccoA Aboli299 Chlamydia trachomatis 518	ORF1100	170509	170793		AE001300		489	100
188295 188023 hypothetical protein AB001298 Chlamydia trachomatis 451 188791 188330 hypothetical protein AB001298 Chlamydia trachomatis 733 190629 190336 putative AB001298 Chlamydia trachomatis 733 210914 211384 putative AB001295 Chlamydia trachomatis 507 235160 234852 Glutamate Aminomutase AB001295 Chlamydia trachomatis 507 237227 236913 putative AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 512 271633 271633 Appothetical protein AB001291 Chlamydia trachomatis 512 271633 271932 Appothetical protein AB001291 Chlamydia trachomatis 512 2775666 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700	ORF1101	177145	177474	AcCoA Carboxylase/Transferase Alpha	AE001299	Chlamydia trachomatis	518	66
188791 188330 hypothetical protein AB001298 Chlamydia trachomatis 733 190629 190336 putative 733 197313 197083 putative 733 210914 211384 putative 7601295 237227 236913 putative AB001295 Chlamydia trachomatis 507 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253701 254789 hypothetical protein AB001293 Chlamydia trachomatis 1860 271633 271932 hypothetical protein AB001291 Chlamydia trachomatis 512 275666 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700, 277931 277931 277931 Abutative AB001291 Chlamydia trachomatis 700,	ORF1102	188295	188023		AE001298	ľ	451	100
190629 190336 putative 190629 190336 putative 197333 197083 putative 701384 700, 70184 700, 70	ORF1103	188791	188330		AE001298	1	733	97
197313 197083 putative AB001295 Chlamydia trachomatis 507 210914 211384 putative AB001295 Chlamydia trachomatis 507 237227 236913 putative AB001293 Chlamydia trachomatis 512 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 1860 271633 271932 hypothetical protein AB001291 Chlamydia trachomatis 512 275666 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700, 277931 278218 putative AB001291 Chlamydia trachomatis 700,	ORF1104	190629	190336	putative				
210914 211384 putative AB001295 Chlamydia trachomatis 507 237227 236913 putative AB001295 Chlamydia trachomatis 512 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 318 253701 254789 hypothetical protein AB001293 Chlamydia trachomatis 1860 271633 271633 271932 Disulfide bond AB001291 Chlamydia trachomatis 700, 277566 275070 Oxidoreductase AB001291 Chlamydia trachomatis 700,	ORF1105	197313	197083	putative				
235160 234852 Glutamate Aminomutase AB001295 Chlamydia trachomatis 507 237227 236913 putative AB001293 Chlamydia trachomatis 512 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AE001293 Chlamydia trachomatis 318 253701 254789 hypothetical protein AE001293 Chlamydia trachomatis 1860 271633 271932 Disulfide bond AE001291 Chlamydia trachomatis 512 277566 276070 Oxidoreductase AE001291 Chlamydia trachomatis 700.	ORF1106	210914	211384	putative				
237227 236913 putative AB001293 Chlamydia trachomatis 512 249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 1860 253701 254789 hypothetical protein AB001291 Chlamydia trachomatis 1860 271633 271932 hypothetical protein AB001291 Chlamydia trachomatis 512 275666 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700, 277931 277931 278218 putative 700,	ORF1107	235160	234852	Glutamate Aminomutase	AE001295	Chlamydia trachomatis	507	97
249733 249446 Oligopeptide Permease AB001293 Chlamydia trachomatis 512 253493 253158 hypothetical protein AB001293 Chlamydia trachomatis 318 253701 254789 hypothetical protein AB001293 Chlamydia trachomatis 1860 271633 271633 Disulfide bond AB001291 Chlamydia trachomatis 512 2775666 276070 Oxidoreductase AB001291 Chlamydia trachomatis 700, 277931 277931 278218 putative 700,	ORF1108	237227	236913	putative				
253493 253158 hypothetical protein AE001293 Chlamydia trachomatis 318 253701 254789 hypothetical protein AE001293 Chlamydia trachomatis 1860 271633 271633 276070 Disulfide bond AE001291 Chlamydia trachomatis 512 277931 278218 putative Putative 700,	ORF1109	249733	249446	Oligopeptide Permease	AE001293		512	100
253701 254789 hypothetical protein AE001293 Chlamydia trachomatis 1860 271633 271633 271932 hypothetical protein AE001291 Chlamydia trachomatis 512 275666 276070 Oxidoreductase AE001291 Chlamydia trachomatis 700, 277931 278218 putative 700,	ORF1110	253493	253158		AE001293		318	63
271633 271932 hypothetical protein AE001291 Chlamydia trachomatis 512 275666 276070 Disulfide bond AE001291 Chlamydia trachomatis 700. 277931 277931 278218 putative 700.	ORF1111	253701	254789		AE001293	ŀ	1860	66
275666 276070 Disulfide bond Oxidoreductase AB001291 Chlamydia trachomatis 700, 277931 277931 278218 putative	ORF1112	271633	271932		AE001291	Chlamydia trachomatis	512	100
277931 278218	ORF1113	275666	276070	Disulfide bond Oxidoreductase	AE001291	1	, 007	66
	ORF1114	277931	278218	putative				

ORF	begin	stop	Homology	4			Į,
			hypothetical protein	AE001290	Chlamydia trachomatic	And	P 0
ORF1115	282741	282481	4	AE001230	Ciilailyula Liachomails	477	y 2
ORF1116	293178	293489	Phospholipase D Endonuclease Superfamily	AE001289	Chlamydia trachomatis	433	95
ORF1117	303155	303469	putative				
ORF1118	309297	308965	hypothetical protein	AE001287	Chlamydia trachomatis	422	95
ORF1119	312219	312536	putative				
ORF1120	312853	312602	hypothetical protein	AE001287	Chlamydia trachomatis	338	66
ORF1121	313167	312772	hypothetical protein	AE001287	Chlamydia trachomatis	616	98
ORF1122	320224	320598	hypothetical protein	AE001286	Chlamydia trachomatis	628	98
ORF1123	340249	340503	Oligopeptidase	AE001285	Chlamydia trachomatis	444	100
ORF1124	352839	353324	hypothetical protein	AE001284	Chlamydia trachomatis	751	98
ORF1125	373475	373699	Phopholipase D Superfamily [leader (33) peptide}	AE001282	Chlamydia trachomatis	378	100
ORF1126	377316	377756	hypothetical protein	AE001282	Chlamydia trachomatis	764	99
ORF1127	379268	379657	hypothetical protein	AE001282	Chlamydia trachomatis	535	100
ORF1128	395098	394823	putative				
ORF1129	401594	401142	Flagellar Secretion Protein	AE001280	Chlamydia trachomatis	698	100
ORF1130	410045	410539	hypothetical protein	AE001279	Chlamydia trachomatis	767	100
ORF1131	411425	411658	Coproporphyrinogen III Oxidase	AE001279	Chlamydia trachomatis	399	66
ORF1132	414937	414416	putative				

18	100	100	86	94	96				100	96	95		100	89	81	100	66	100
Score	206	610	1010	649	538				370	483	864		964	360	242	127	683	534
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis				Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis
GI.	AE001278	AE001278	AE001278	AE001277	AE001277				AE001361	AE001360	AE001359		AE001358	AE001358	AE001356	AE001356	AE001354	AE001353
Homology	Glycogen Hydrolase (debranching)	hypothetical protein	hypothetical protein	hypothetical protein	L31 Ribosomal Protein	putative	putative	putative	Putative Outer Membrane Protein I	Membrane Thiol Protease	Low Calcium Response Protein H	putative	ABC transporter permease	hypothetical protein				
stop	423212	428183	429451	442799	444041	443166	446155	468262	472108	488337	497694	500202	509561	511264	526848	531863	557224	564537
begin	422889	427842	428732	442557	443628	443678	445901	467981	471869	488032	497179	500474	508968	510845	526525	531318	556826	564971
ORF	ORF1133	ORF1134	ORF1135	ORF1136	ORF1137	ORF1138	ORF1139	ORF1140	ORF1141	ORF1142	ORF1143	ORF1144	ORF1145	ORF1146	ORF1147	ORF1148	ORF1149	ORF1150

1%	53	100	66	97	100	98		66					66		Τ	86				98		96
Score	220	925	441	273	442	1176		434					324			864				606		661
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis					Chlamydia trachomatis			Chlamydia trachomatis				Chlamydia trachomatis		Chlamydia trachomatis
Ð	AE001353	AE001353	AE001353	AE001352	AE001351	AE001350		AE001349					AE001345			AE001343				AE001336		AE001335
Homology	Glycerol-3-P Acyltransferase	Insulinase family/Protease III	hypothetical protein	General Stress Protein	hypothetical protein	hypothetical protein	putative	hydrolase/phosphatase homolog	putative	putative	putative	putative	Phenolhydrolase/NADH ubiquinone oxidoreductase	putative	putative	hypothetical protein	putative	putative	putative	(Pseudouridine Synthase)	putative	hypothetical protein
stop	567232	570890	571332	575801	590650	598593	606626	607786	610391	633353	637482	649924	652562	655325	660810	677057	679253	732210	742383	758782	760521	770894
begin	566963	570351	571072	576025	590363	597868	688909	608031	610110	632703	637213	650517	652317	654753	661118	677596	679528	732536	742069	759318	760282	771313
ORF	ORF1151	ORF1152	ORF1153	ORF1154	ORF1155	ORF1156	ORF1157	ORF1158	ORF1159	ORF1160	ORF1161	ORF1162	ORF1163	ORF1164	ORF1165	ORF1166	ORF1167	ORF1168	ORF1169	ORF1170	ORF1171	ORF1172

		-,-	_	_																	•	
%i	66					86	66		95				97		66	100	98		97		86	
Score	520					268	747		551				195		410	593	542		467		647	
Species	Chlamydia trachomatis					Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis				Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis		Chlamydia trachomatis	
ΩĪ	AE001335					AE001327	AE001327		AE001326				AE001324		AE001322	AE001322	AE001321		AE001320		AE001320	
Homology	hypothetical protein	putative	putative	putative	putative	hypothetical protein	hypothetical protein	putative	hypothetical protein	putative	putative	putative	Myristoyl GlcNac Deacetylase	putative	hypothetical protein	hypothetical protein	hypothetical protein	putative	2-Component Sensor	putative	Phosphoglycolate Phosphatase	putative
stop	772408	788457	815967	846914	868054	829528	876915	884312	891467	900417	902269	907783	912567	935741	946692	952783	965873	968765	970731	972404	973508	998404
begin	772115	788137	816302	846606	867803	875386	876445	884548	891859	900770	902553	908046	912313	935451	946961	953193	966199	969298	971009	972162	973119	998649
ORF	ORF1173	ORF1174	ORF1175	ORF1176	ORF1177	ORF1178	ORF1179	ORF1180	ORF1181	ORF1182	ORF1183	ORF1184	ORF1185	ORF1186	ORF1187	ORF1188	ORF1189	ORF1190	ORF1191	ORF1192	ORF1193	ORF1194

ORF	begin	stop	Homology	Ð	Species	Score	H
ORF1195	1004280	1003882	hypothetical protein	AE001317	Chlamydia trachomatis	571	66
ORF1196	1010200	1009532	hypothetical protein	AE001317	AE001317 Chlamydia trachomatis	1132	66
ORF1197	1029174	1029482	putative				

TABLE 2

2 501 208 501 3 3276 505 3153 4 5068 3242 5062 5 6400 5126 6400 6 7977 6619 7977 7 8582 8082 8582 8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21	SEQ ID NO	begin	stop	preferred start
4 5068 3242 5062 5 6400 5126 6400 6 7977 6619 7977 7 8582 8082 8582 8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 <	2	501	208	501
5 6400 5126 6400 6 7977 6619 7977 7 8582 8082 8582 8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795	3	3276	505	3153
6 7977 6619 7977 7 8582 8082 8582 8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180	4	5068	3242	5062
7 8582 8082 8582 8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383	5	6400	5126	6400
8 8995 8591 8995 9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 </td <td>6</td> <td>7977</td> <td>6619</td> <td>7977</td>	6	7977	6619	7977
9 9440 8979 9440 10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729	7	8582	8082	8582
10 9828 10430 9828 11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640	8	8995	8591	8995
11 10367 11254 10430 12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 <	9	9440	8979	9440
12 11245 11916 11245 13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 <	10	9828	10430	9828
13 12068 13324 12068 14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	11	10367	11254	10430
14 13532 14413 13538 15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	12	11245	11916	11245
15 14807 15019 14807 16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	13	12068	13324	12068
16 14932 15969 14977 17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	14	13532	14413	13538
17 15995 16501 16004 18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	15	14807	15019	14807
18 16467 16138 16377 19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	16	14932	15969	14977
19 18190 17417 18178 20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	17	15995	16501	16004
20 20521 18437 20518 21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	18	16467	16138	16377
21 22202 20814 22166 22 22602 22153 22509 23 22804 22478 22795 24 23183 22824 23180 25 23394 23110 23394 26 24569 23394 24569 27 26383 24641 26383 28 26640 27710 26640 29 28780 27725 28729 30 29957 28740 29957 31 30721 30032 30628 32 31281 30520 31254	19	18190	17417	18178
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35	33901	33314	33874
36	34116	35027	34116
37	34988	35359	35027
38	35167	35919	35377
39	35923	36996	36031
40	37810	37013	37765
41	38207	39085	38252
42	39151	39927	39157
43	39923	40756	39959
44	40760	42007	40772
45	42175	43116	42229
46	42999	43802	43128
47	44211	45227	44217
48	46072	45275	46066
49	46340	45975	46331
50	46895	46506	46865
51	47955	46882	47955
52	48585	48178	48558
53	50072	48630	50012
54	50710	50099	50692
55	52439	50925	52430
56	53484	52348	53478
57	54536	53466	54536
58	55086	54595	55104
59	56350	55031	56350
60	55659	56084	55722
61	56847	58235	56931
62	58423	59181	58423
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64	60188	61483	60191
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66	62500	63141	62518
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69	64285	64656	64285
70	64944	64609	64938
71	65347	67269	65347
72	67656	68873	67815
73	68877	69233	68892
74	69212	69721	69323
75	69958	70455	69970
76	70701	71006	70725
77	73191	71086	73185
78	74900	73497	74891
79	75463	74876	75463
80	77124	75502	77124
81	77000	77299	77012
82	78095	77145	78095
83	79065	78154	79065
84	81971	79878	81965
85	82639	83271	82642
86	83792	84850	83921
87	84876	86921	84888
88	88650	87313	88383
89	87440	87805	87458
90	88400	88747	88409
91	88717	89265	88729
92	89355	89732	89355
93	89735	91447	89735
94	91749	91435	91749
95	92392	91745	92323
96	93138	92344	92874
97	94134	93361	93945
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107	105259	105894	105271
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109	108665	108955	108683
110	109459	109013	109456
111	110366	109704	110363
112	111330	112520	111330
113	112915	113463	112918
114	113566	113994	113566
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117	115362	115676	115380
118	116022	119795	116040
119	119823	124010	119823
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123	126328	126930	126331
124	127138	127785	127147
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126	129720	131033	129720
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128	131834	133156	131852
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130	133625	133999	133628
131	133861	134508	133948
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140	146413	147078	146425
141	147140	148075	147152
142	148115	148549	148115
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145	149187	149708	149187
146	149712	150911	149769
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151	155276	153984	155228
152	156544	155231	156544
153	156806	157525	156809
154	157489	158955	157534
155	159104	159961	159104
156	159916	161220	159916
157	161183	161593	161228
158	162354	161623	162354
159	163013	162363	163013
160	163941	162994	163941
161	165505	164474	165505
162	166686	166093	166686
163	168171	166729	168171
164	169249	168848	169189
165	169586	170431	169607
166	170780	171334	170783
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168	172309	172722	172309
169	173048	174496	173048

SEQ ID NO	begin	stop	preferred start
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174	178084	180021	178084
175	180704	180048	180635
176	181398	180631	181398
177	182594	181398	182594
178	182895	183656	182895
179	183665	184786	183665
180	186007	184796	186007
181	186848	186000	186791
182	187270	186749	187240
183	187426	187809	187429
184	189481	188798	189442
185	189693	190352	189693
186	190235	190510	190280
187	190785	191786	190824
188	191790	192464	191811
189	192392	193183	192500
190	193254	194630	193263
191	195046	194690	195037
192	195184	197031	195193
193	197018	197635	197024
194	197762	198208	197669
195	198963	197668	198954
196	199957	198962	199945
197	200327	199941	200306
198	200685	200266	200598
199	200962	200585	200962
200	201169	202377	201184
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207	209471	210214	209471
208	210586	210816	210586
209	211332	210883	211293
210	212978	211374	212972
211	214134	212875	214134
212	214710	214168	214701
213	215143	214754	215128
214	216705	215236	216705
215	217917	216892	217911
216	217088	217441	217202
217	218364	218702	218364
218	218695	219009	218785
219	219179	219748	219260
220	219891	220430	219912
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222	221137	221541	221176
223	221601	222092	221616
224	222472	223290	222487
225	223423	223818	223423
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227	225749	225174	225749
228	225334	225549	225328
229	226654	225749	226654
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231	227646	227161	227646
232	228457	227750	228439
233	230001	228607	229854
234	231074	230151	231062
235	231348	233006	231366
236	233059	233829	233059
237	233801	234265	233801

SEQ ID NO	begin	Stop	preferred start
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240	236314	238209	236314
241	238164	238769	238185
242	238769	240061	238769
243	242022	240313	242022
244	242846	241941	242846
245	244480	242798	244456
246	245897	244479	245891
247	246877	245924	246829
248	247731	246985	247725
249	248585	247743	248573
250	249420	248569	249411
251	250383	249766	250383
252	251186	250545	251174
253	252111	251095	252099
254	253088	252066	253088
255	255153	256718	255153
256	256762	257844	256774
257	257911	258690	257962
258	258780	259187	258840
259	259193	261604	259193
260	261622	264129	261622
261	264125	264742	264134
262	264741	265628	264759
263	266416	265631	266416
264	266938	266426	267946
265	267961	266942	267946
266	268320	268066	268299
267	268510	268205	268510
268	270116	268500	270116
269	270892	270095	270856
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271	272219	272932	272243

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274	274821	275666	274953
275	277689	276103	277689
276	278268	278816	278298
277	279771	279013	279870
278	280777	279767	280762
279	281603	281295	281576
280	282104	281787	282086
281	284335	282794	284320
282	284460	284795	284550
283	284817	285674	284844
284	285637	286137	285670
285	286357	286677	286399
286	286681	287898	286852
287	288127	289227	288358
288	289744	290679	289744
289	290828	291535	291206
290	291514	292230	291514
291	292326	293048	292350
292	293330	294853	293525
293	295684	295010	295684
294	296336	295692	296294
295	297238	296243	297199
296	297791	298735	297791
297	298905	300458	298920
298	302152	300527	302131
299	304917	302071	304872
300	306157	304973	306142
301	306494	306111	306461
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303	308773	306977	308758
304	309881	309276	309869
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311	316507	315665	316507
312	317284	316529	317284
313	317592	317338	317592
314	318470	317499	318416
315	317599	317874	317599
316	318947	318477	318887
317	319342	320142	319342
318	320544	321497	320682
319	321485	321937	321497
320	321901	322362	321943
321	322301	323140	322325
322	323144	324913	323177
323	325621	324977	325621
324	326268	325621	326262
325	326469	327203	326469
326	327281	328150	327302
327	328605	328204	328602
328	329066	328734	329066
329	329663	329292	329648
330	330666	329608	330663
331	331161	330670	331161
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333	332404	331721	332404
334	332779	333021	332779
335	333005	333589	333149
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341	336334	335903	336229
342	337378	336338	337378
343	339947	337347	339947
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345	341783	342022	341786
346	342249	342470	342249
347	342597	343370	342597
348	343361	344032	343379
349	343956	344225	343962
350	344357	345142	344357
351	345934	345161	345934
352	347102	346080	347102
353	347113	347940	347119
354	350164	348146	350113
355	350423	351283	350426
356	352207	351314	352207
357	352727	352245	352703
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359	354218	353670	354215
360	354721	354140	354721
361	354966	356672	354966
362	356700	357377	356700
363	357326	358093	357500
364	358035	360743	358035
365	360753	361121	360753
366	361162	361884	361162
367	361826	362746	361826
368	363859	362816	363859
369	364116	365195	364116
370	365198	365587	365198
371	365479	367320	365614
372	367341	368603	367341
373	368644	369081	368644

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376	371203	372816	371209
377	373119	373529	373152
378	373614	374204	373776
379	374736	374224	374703
380	376391	374703	376382
381	377062	376748	377038
382	377853	378737	377871
383	378626	379048	378710
384	379017	379403	379038
385	380009	379641	379967
386	380187	381470	380187
387	381473	382567	381473
388	382704	383702	382728
389	383945	383655	383921
390	385217	383949	385211
391	385507	385178	385507
392	386845	385706	386842
393	386127	386627	386232
394	387372	386872	387351
395	387823	387338	387823
396	388250	387816	388106
397	389169	388237	389169
398	389955	389173	390087
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401	392410	393996	392413
402	394170	395354	394185
403	395309	395992	395354
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405	397507	396542	397498
406	398753	397401	398747
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413	403788	403276	403785
414	405165	405920	405165
415	407049	405955	407049
416	409773	407056	409764
417	410532	411416	410532
418	411707	413410	411722
419	413433	412606	413334
420	413404	413952	413449
421	413841	415112	413991
422	414379	413978	414220
423	416664	415177	416646
424	417456	416740	417456
425	418053	417721	418044
426	418603	418031	418582
427	419531	418647	419531
428	420190	419672	420190
429	421171	420245	421171
430	421988	421518	421988
431	422486	423043	422492
432	423226	425079	423295
433	426054	425146	426021
434	426985	426245	426967
435	427248	427817	427248
436	429560	429886	429623
437	430360	429857	430360
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439	430933	431787	430966
440	431658	431987	431688
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SEQ ID NO	begin	stop	preferred star
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448	439705	439358	439705
449	441042	439699	441042
450	441911	441042	441911
451	442593	441898	442584
452	444505	446388	444505
453	448068	446452	448029
454	449575	447932	449575
455	450546	451076	450576
456	451623	451144	451401
457	452593	451517	452575
458	453195	452632	453174
459	453567	454868	453567
460	455430	454972	455403
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466	461624	461196	461624
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468	463758	462895	463749
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471	467420	466113	467414
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473	469280	468906	469226
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SEQ ID NO	begin	stop	preferred star
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481	476527	475613	476509
482	478640	476517	478640
483	479084	478665	479078
484	479723	479088	479720
485	480012	479668	479898
486	481466	479895	481412
487	481732	481496	481732
488	481864	483429	481870
489	483402	484964	483402
490	484898	487864	484970
491	485725	485222	485593
492	488204	489247	488321
493	488571	488233	488562
494	489440	490456	489473
495	492765	490507	492690
496	492357	492893	492654
497	493744	492737	493723
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501	495174	494872	495174
502	495687	496634	495732
503	496295	497176	496523
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516	509600	510691	509600
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518	511547	512185	511547
519	512382	513092	512385
520	514287	513055	514269
521	514789	515244	514792
522	514994	515269	515027
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526	517927	517400	517927
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528	518403	518822	518412
529	518923	519516	518923
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533	523495	522143	523483
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552	540906	541805	541002
553	543255	541825	543222
554	544133	543222	544115
555	544565	544179	544532
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829	815213	814314	815207
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832	816116	817456	816170
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857	836602	837264	836617
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862	841659	842459	841686
863	842523	843068	842541
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865	843239	846196	843335
866	844137	843802	844077
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868	850123	848150	850099
869	851645	850230	851504
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873	856240	855437	856240
874	857183	856233	857006
875	859439	857451	859430
876	859946	859587	859916
877	859642	860640	859660
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881	863930	863571	863927
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889	871299	872582	871359
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891	872773	873915	872773
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SEQ ID NO	begin	Stop	preferred start
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933	912399	912629	912399
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936	913691	914485	913691
937	914516	915136	914522
938	915144	915467	915162
939	915629	916633	915629
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941	916965	917627	916965
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946	919723	920157	919723
947	920184	920840	920184
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957	924443	924937	924443
958	924933	925364	924933
959	925390	926760	925390
960	926819	927184	926819
961	927209	927604	927209
962	927577	928155	927577
963	928100	928759	928127
964	929222	930244	929243
965	930222	930656	930258
966	930608	931078	930665
967	931367	931666	931406
968	931549	931959	931558
969	932070	932579	932070
970	932602	933201	932602
971	933319	933621	933319
972	933522	933785	933522
973	934546	933848	934546
974	936377	934539	936377
975	938081	936666	938081
976	938538	939098	938595
977	939329	940933	939506
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979	942082	944685	942082
980	944634	945287	944673
981	945287	946294	945287
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991	954426	955157	954429
992	955754	957940	955766
993	957837	959312	957867
994	959299	961050	959317
995	961562	961053	961562
996	962575	961487	962545
997	961979	961584	961979
998	964914	962545	964914
999	964941	965708	964956
1000	967023	966193	966984
1001	967444	968061	967459
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1003	970685	969528	970685
1004	971806	971024	971785
1005	973053	972388	973026
1006	974546	973746	974546
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1009	976520	976254	976511
1010	976588	976899	976588
1011	976886	977635	976934
1012	977661	977933	977682
1013	977918	978433	977933
1014	978619	978984	978619
1015	978933	979331	978987
1016	981197	979389	981197
1017	979711	980112	979753
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1019	982321	983598	982321

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1023 986693 986046 986693 1024 987607 986693 987607 1025 988119 987616 987942 1026 988253 987936 988247 1027 988831 989163 988834 1028 989693 993442 989693 1030 993835 993416 993754 1031 993882 994262 993906 1032 994226 995656 994259 1033 996036 996611 996036 1034 996885 998267 996885 1035 998962 999225 998962 1036 999375 1001033 999393 1037 1001211 1001516 1001214 1038 1001392 1001664 1001443 1039 1003721 1001823 1003721 1040 1004459 1004845 1004990 1042 1005391 1007496 1005391				985381
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1026 988253 987936 988247 1027 988831 989163 988834 1028 989693 993442 989693 1029 993408 993785 993408 1030 993835 993416 993754 1031 993882 994262 993906 1032 994226 995656 994259 1033 996036 996611 996036 1034 996885 998267 996885 1035 998962 999225 998962 1036 999375 1001033 999393 1037 1001211 1001516 1001214 1038 1001392 1001664 1001443 1039 1003721 1001823 1003721 1040 1004459 1004845 1004459 1041 1004990 1005382 1004990 1042 1005391 1007496 1005391 1043 1007486 1007821 1007453 <td>1024</td> <td>987607</td> <td>986693</td> <td>987607</td>	1024	987607	986693	987607
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1028 989693 993442 989693 1029 993408 993785 993408 1030 993835 993416 993754 1031 993882 994262 993906 1032 994226 995656 994259 1033 996036 996611 996036 1034 996885 998267 996885 1035 998962 999225 998962 1036 999375 1001033 999393 1037 1001211 1001516 1001214 1038 1001392 1001664 1001443 1039 1003721 1001823 1003721 1040 1004459 1004845 1004459 1041 1004990 1005382 1004990 1042 1005391 1007496 1005391 1043 1007486 1007821 1007453 1044 1007802 1008698 1007841 1045 1009426 1009121 10094	1026	988253	987936	988247
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1048 1012042 1012635 1012057 1049 1012593 1012862 1012593 1050 1012811 1013440 1012829 1051 1013456 1014055 1013468 1052 1013977 1014489 1013977	1046	1010534	1012054	1010534
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1051 1013456 1014055 1013468 1052 1013977 1014489 1013977	1049	1012593	1012862	1012593
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	1051	1013456	1014055	1013468
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1056	1017766	1017245	1017658
1057	1018911	1017243	1017638
1058	1019111	1017916	1018893
1059	1020199	1019831	
1060	1021007	1019831	1020196
1061			1020992
	1021569	1021075	1021557
1062	1022411	1022097	1022402
1063	1023344	1023667	1023344
1064	1023701	1023949	1023701
1065	1023976	1024776	1023976
1066	1024704	1025045	1024704
1067	1025881	1024967	1025845
1068	1026546	1025839	1026546
1069	1027379	1026546	1027373
1070	1030604	1027929	1030328
1071	1033252	1030508	1033249
1072	1031733	1032086	1031823
1073	1037037	1033456	1037016
1074	1035674	1035910	1035674
1075	1036175	1036507	1036268
1076	68(comp)	1036967	38
1077	16591	16989	16597
1078	31779	31408	31764
1079	56502	56834	56520
1080	56686	56913	56686
1081	64748	65074	64790
1082	73482	73195	73482
1083	78482	78736	78506
1084	79803	79411	79773
1085	82333	81959	82333
1086	87313	86999	87523
1087	109929	109456	109716

SEQ ID NO	begin	stop	preferred start
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1089	112069	111734	111988
1090	112666	112911	112666
1091	114017	113715	113978
1092	120757	120464	120757
1093	125133	125522	125133
1094	131888	131604	131837
1095	144164	144427	144191
1096	150698	150369	150635
1097	164385	163948	164385
1098	165690	166115	165408
1099	168742	168425	168742
1100	170509	170793	170509
1101	177145	177474	177145
1102	188295	188023	188295
1103	188791	188330	188791
1104	190629	190336	190626
1105	197313	197083	197307
1106	210914	211384	210956
1107	235160	234852	235160
1108	237227	236913	237188
1109	249733	249446	249904
1110	253493	253158	253493
1111	253701	254789	253701
1112	271633	271932	271633
1113	275666	276070	275666
1114	277931	278218	277976
1115	282741	282481	282738
1116	293178	293489	293181
1117	303155	303469	303185
1118	309297	308965	309297
1119	312219	312536	312246
1120	312853	312602	312844
1121	313167	312772	313167

SEQ ID NO	begin	stop	preferred start
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1123	340249	340503	340249
1124	352839	353324	352839
1125	373475	373699	373475
1126	377316	377756	377316
1127	379268	379657	379268
1128	395098	394823	395077
1129	401594	401142	401594
1130	410045	410539	410045
1131	411425	411658	411425
1132	414937	414416	414937
1133	422889	423212	422964
1134	427842	428183	427842
1135	428732	429451	428732
1136	442557	442799	442524
1137	443628	444041	443628
1138	443678	443166	443678
1139	445901	446155	445901
1140	467981	468262	468023
1141	471869	472108	471869
1142	488032	488337	488044
1143	497179	497694	497101
1144	500474	500202	500471
1145	508968	509561	508968
1146	510845	511264	510845
1147	526525	526848	526525
1148	531318	531863	531444
1149	556826	557224	556826
1150	564971	564537	564971
1151	566963	567232	566963
1152	570351	570890	570351
1153	571072	571332	571072
1154	576025	575801	576025
1155	590363	590650	590363

SEQ ID NO	begin	stop	preferred start
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1158	608031	607786	608031
1159	610110	610391	610143
1160	632703	633353	632703
1161	637213	637482	637255
1162	650517	649924	650517
1163	652317	652562	652317
1164	654753	655325	654753
1165	661118	660810	661118
1166	677596	677057	677578
1167	679528	679253	679477
1168	732536	732210	732536
1169	742069	742383	742069
1170	759318	758782	759318
1171	760282	760521	760282
1172	771313	770894	771391
1173	772115	772408	772115
1174	788137	788457	788137
1175	816302	815967	816302
1176	846606	846914	846612
1177	867803	868054	867806
1178	875386	875658	875395
1179	876445	876915	876445
1180	884548	884312	884548
1181	891859	891467	891859
1182	900770	900417	900728
1183	902553	902269	902529
1184	908046	907783	908007
1185	912313	912567	912313
1186	935451	935741	935451
1187	946961	946692	946940
1188	953193	952783	953145
1189	966199	965873	966184

SEQ ID NO	begin	stop	preferred start
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1191	971009	970731	971009
1192	972162	972404	972165
1193	973119	973508	973119
1194	998649	998404	998625
1195	1004280	1003882	1004280
1196	1010200	1009532	1010200
1197	1029174	1029482	1029180

TABLE 4

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
2	1199	1198	3591	3590
3	1201	1200	3593	3592
4	1203	1202	3595	3594
5	1205	1204	3597	3596
6	1207	1206	3599	3598
7	1209	1208	3601	3600
8	1211	1210	3603	3602
9	1213	1212	3605	3604
10	1215	1214	3607	3606
11	1217	1216	3609	3608
12	1219	1218	3611	3610
13	1221	1220	3613	3612
14	1223	1222	3615	3614
15	1225	1224	3617	3616
16	1227	1226	3619	3618
17	1229	1228	3621	3620
18	1231	1230	3623	3622
19	1233	1232	3625	3624
20	1235	1234	3627	3626
21	1237	1236	3629	3628
22	1239	1238	3631	3630
23	1241	1240	3633	3632
24	1243	1242	3635	3634
25	1245	1244	3637	3636
26	1247	1246	3639	3638
27	1249	1248	3641	3640
28	1251	1250	3643	3642
29	1253	1252	3645	3644
30	1255	1254	3647	3646
31	1257	1256	3649	3648
32	1259	1258	3651	3650
33	1261	1260	3653	3652

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
34	1263	1262	3655	3654
35	1265	1264	3657	3656
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37	1269	1268	3661	3660
38	1271	1270	3663	3662
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43	1281	1280	3673	3672
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46	1287	1286	3679	3678
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48	1291	1290	3683	3682
49	1293	1292	3685	3684
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51	1297	1296	3689	3688
52	1299	1298	3691	3690
53	1301	1300	3693	3692
54	1303	1302	3695	3694
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56	1307	1306	3699	3698
57	1309	1308	3701	3700
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59	1313	1312	3705	3704
60	1315	1314	3707	3706
61	1317	1316	3709	3708
62	1319	1318	3711	3710
63	1321	1320	3713	3712
64	1323	1322	3715	3714
65	1325	1324	3717	3716
66	1327	1326	3719	3718
67	1329	1328	3721	3720

	194					
ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp		
68	1331	1330	3723	3722		
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70	1335	1334	3727	3726		
71	1337	1336	3729	3728		
72	1339	1338	3731	3730		
73	1341	1340	3733	3732		
74	1343	1342	3735	3734		
75	1345	1344	3737	3736		
76	1347	1346	3739	3738		
77	1349	1348	3741	3740		
78	1351	1350	3743	3742		
79	1353	1352	3745	3744		
80	1355	1354	3747	3746		
81	1357	1356	3749	3748		
82	1359	1358	3751	3750		
83	1361	1360	3753	3752		
84	1363	1362	3755	3754		
85	1365	1364	3757	3756		
86	1367	1366	3759	3758		
87	1369	1368	3761	3760		
88	1371	1370	3763	3762		
89	1373	1372	3765	3764		
90	1375	1374	3767	3766		
91	1377	1376	3769	3768		
92	1379	1378	3771	3770		
93	1381	1380	3773	3772		
94	1383	1382	3775	3774		
95	1385	1384	3777	3776		
96	1387	1386	3779	3778		
97	1389	1388	3781	3780		
98	1391	1390	3783	3782		
99	1393	1392	3785	3784		
100	1395	1394	3787	3786		
101	1397	1396	3789	3788		
		·				

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
102	1399	1398	3791	3790
103	1401	1400	3793	3792
104	1403	1402	3795	3794
105	1405	1404	3797	3796
106	1407	1406	3799	3798
107	1409	1408	3801	3800
108	1411	1410	3803	3802
109	1413	1412	3805	3804
110	1415	1414	3807	3806
111	1417	1416	3809	3808
112	1419	1418	3811	3810
113	1421	1420	3813	3812
114	1423	1422	3815	3814
115	1425	1424	3817	3816
116	1427	1426	3819	3818
117	1429	1428	3821	3820
118	1431	1430	3823	3822
119	1433	1432	3825	3824
120	1435	1434	3827	3826
121	1437	1436	3829	3828
122	1439	1438	3831	3830
123	1441	1440	3833	3832
124	1443	1442	3835	3834
125	1445	1444	3837	3836
126	1447	1446	3839	3838
127	1449	1448	3841	3840
128	1451	1450	3843	3842
129	1453	1452	3845	3844
130	1455	1454	3847	3846
131	1457	1456	3849	3848
132	1459	1458	3851	3850
∠133	1461	1460	3853	3852
134	1463	1462	3855	3854
135	1465	1464	3857	3856

136 1467 1466 3859 3858 137 1469 1468 3861 3860 138 1471 1470 3863 3862 139 1473 1472 3865 3864 140 1475 1474 3867 3866 141 1477 1476 3869 3868 142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3889 </th <th>ORFGenset</th> <th>ORFoligosFd</th> <th>ORFoligosFp</th> <th>ORFoligosBd</th> <th>ORFoligosBp</th>	ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
138 1471 1470 3863 3862 139 1473 1472 3865 3864 140 1475 1474 3867 3866 141 1477 1476 3869 3868 142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 </td <td>136</td> <td>1467</td> <td>1466</td> <td>3859</td> <td></td>	136	1467	1466	3859	
139 1473 1472 3865 3864 140 1475 1474 3867 3866 141 1477 1476 3869 3868 142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 </td <td>137</td> <td>1469</td> <td>1468</td> <td>3861</td> <td>3860</td>	137	1469	1468	3861	3860
140 1475 1474 3867 3866 141 1477 1476 3869 3868 142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 </td <td>138</td> <td>1471</td> <td>1470</td> <td>3863</td> <td>3862</td>	138	1471	1470	3863	3862
141 1477 1476 3869 3868 142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 </td <td>139</td> <td>1473</td> <td>1472</td> <td>3865</td> <td>3864</td>	139	1473	1472	3865	3864
142 1479 1478 3871 3870 143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 </td <td>140</td> <td>1475</td> <td>1474</td> <td>3867</td> <td>3866</td>	140	1475	1474	3867	3866
143 1481 1480 3873 3872 144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 </td <td>141</td> <td>1477</td> <td>1476</td> <td>3869</td> <td>3868</td>	141	1477	1476	3869	3868
144 1483 1482 3875 3874 145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 </td <td>142</td> <td>1479</td> <td>1478</td> <td>3871</td> <td>3870</td>	142	1479	1478	3871	3870
145 1485 1484 3877 3876 146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 </td <td>143</td> <td>1481</td> <td>1480</td> <td>3873</td> <td>3872</td>	143	1481	1480	3873	3872
146 1487 1486 3879 3878 147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 </td <td>144</td> <td>1483</td> <td>1482</td> <td>3875</td> <td>3874</td>	144	1483	1482	3875	3874
147 1489 1488 3881 3880 148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 </td <td>145</td> <td>1485</td> <td>1484</td> <td>3877</td> <td>3876</td>	145	1485	1484	3877	3876
148 1491 1490 3883 3882 149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 </td <td>146</td> <td>1487</td> <td>1486</td> <td>3879</td> <td>3878</td>	146	1487	1486	3879	3878
149 1493 1492 3885 3884 150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166	147	1489	1488	3881	3880
150 1495 1494 3887 3886 151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 </td <td>148</td> <td>1491</td> <td>1490</td> <td>3883</td> <td>3882</td>	148	1491	1490	3883	3882
151 1497 1496 3889 3888 152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 </td <td>149</td> <td>1493</td> <td>1492</td> <td>3885</td> <td>3884</td>	149	1493	1492	3885	3884
152 1499 1498 3891 3890 153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 </td <td>150</td> <td>1495</td> <td>1494</td> <td>3887</td> <td>3886</td>	150	1495	1494	3887	3886
153 1501 1500 3893 3892 154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922 </td <td>151</td> <td>1497</td> <td>1496</td> <td>3889</td> <td>3888</td>	151	1497	1496	3889	3888
154 1503 1502 3895 3894 155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	152	1499	1498	3891	3890
155 1505 1504 3897 3896 156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	153	1501	1500	3893	3892
156 1507 1506 3899 3898 157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	154	1503	1502	3895	3894
157 1509 1508 3901 3900 158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	155	1505	1504	3897	3896
158 1511 1510 3903 3902 159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	156	1507	1506	3899	3898
159 1513 1512 3905 3904 160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	157	1509	1508	3901	3900
160 1515 1514 3907 3906 161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	158	1511	1510	3903	3902
161 1517 1516 3909 3908 162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	159	1513	1512	3905	3904
162 1519 1518 3911 3910 163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	160	1515	1514	3907	3906
163 1521 1520 3913 3912 164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	161	1517	1516	3909	3908
164 1523 1522 3915 3914 165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	162	1519	1518	3911	3910
165 1525 1524 3917 3916 166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	163	1521	1520	3913	3912
166 1527 1526 3919 3918 167 1529 1528 3921 3920 168 1531 1530 3923 3922	164	1523	1522	3915	3914
167 1529 1528 3921 3920 168 1531 1530 3923 3922	165	1525	1524	3917	3916
168 1531 1530 3923 3922	166	1527	1526	3919	3918
3723 3722	167	1529	1528	3921	3920
169 1533 1532 3925 3924	168	1531	1530	3923	3922
	169	1533	1532	3925	3924

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
170	1535	1534	3927	3926
171	1537	1536	3929	3928
172	1539	1538	3931	3930
173	1541	1540	3933	3932
174	1543	1542	3935	3934
175	1545	1544	3937	3936
176	1547	1546	3939	3938
177	1549	1548	3941	3940
178	1551	1550	3943	3942
179	1553	1552	3945	3944
180	1555	1554	3947	3946
181	1557	1556	3949	. 3948
182	1559	1558	3951	3950
183	1561	1560	3953	3952
184	1563	1562	3955	3954
185	1565	1564	3957	3956
186	1567	1566	3959	3958
187	1569	1568	3961	3960
188	1571	1570	3963	3962
189	1573	1572	3965	3964
190	1575	1574	3967	3966
191	1577	1576	3969	3968
192	1579	1578	3971	3970
193	1581	1580	3973	3972
194	1583	1582	3975	3974
195	1585	1584	3977	3976
196	1587	1586	3979	3978
197	1589	1588	3981	3980
198	1591	1590	3983	3982
199	1593	1592	3985	3984
200	1595	1594	3987	3986
201	1597	1596	3989	3988
202	1599	1598	3991	3990
203	1601	1600	3993	3992

ORFGenset	ORFoligosFd	ORFoligosFp	ODE SPACE	ODE TO
204			ORFoligosBd	ORFoligosBp
	1603	1602	3995	3994
205	1605	1604	3997	3996
206	1607	1606	3999	3998
207	1609	1608	4001	4000
208	1611	1610	4003	4002
209	1613	1612	4005	4004
210	1615	1614	4007	4006
211	1617	1616	4009	4008
212	1619	1618	4011	4010
213	1621	1620	4013	4012
214	1623	1622	4015	4014
215	1625	1624	4017	4016
216	1627	1626	4019	4018
217	1629	1628	4021	4020
218	1631	1630	4023	4022
219	1633	1632	4025	4024
220	1635	1634	4027	4026
221	1637	1636	4029	4028
222	1639	1638	4031	4030
223	1641	1640	4033	4032
224	1643	1642	4035	4034
225	1645	1644	4037	4036
226	1647	1646	4039	4038
227	1649	1648	4041	4040
228	1651	1650	4043	4042
229	1653	1652	4045	4044
230	1655	1654	4047	4046
231	1657	1656	4049	4048
232	1659	1658	4051	4050
233	1661	1660	4053	4052
234	1663	1662	4055	4054
235	1665	1664	4057	4056
236	1667	1666	4059	4058
237	1669	1668	4061	4060
	1007	1000	4001	4000

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
238	1671	1670	4063	4062
239	1673	1672	4065	4064
240	1675	1674	4067	4066
241	1677	1676	4069	4068
242	1679	1678	4071	4070
243	1681	1680	4073	4072
244	1683	1682	4075	4074
245	1685	1684	4077	4076
246	1687	1686	4079	4078
247	1689	1688	4081	4080
248	1691	1690	4083	4082
249	1693	1692	4085	4084
250	1695	1694	4087	4086
251	1697	1696	4089	4088
252	1699	1698	4091	4090
253	1701	1700	4093	4092
254	1703	1702	4095	4094
255	1705	1704	4097	4096
256	1707	1706	4099	4098
257	1709	1708	4101	4100
258	1711	1710	4103	4102
259	1713	1712	4105	4104
260	1715	1714	4107	4106
261	1717	1716	4109	4108
262	1719	1718	4111	4110
263	1721	1720	4113	4112
264	1723	1722	4115	4114
265	1725	1724	4117	4116
266	1727	1726	4119	4118
267	1729	1728	4121	4120
268	1731	1730	4123	4122
269	1733	1732	4125	4124
270	1735	1734	4127	4126
271	1737	1736	4129	4128

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
272	1739	1738	4131	4130
273	1741	1740	4133	4132
274	1743	1742	4135	4134
275	1745	1744	4137	4136
276	1747	1746	4139	4138
277	1749	1748	4141	4140
278	1751	1750	4143	4142
279	1753	1752	4145	4144
280	1755	1754	4147	4146
281	1757	1756	4149	4148
282	1759	1758	4151	4150
283	1761	1760	4153	4152
284	1763	1762	4155	4154
285	1765	1764	4157	4156
286	1767	1766	4159	4158
287	1769	1768	4161	4160
288	1771	1770	4163	4162
289	1773	1772	4165	4164
290	1775	1774	4167	4166
291	1777	1776	4169	4168
292	1779	1778	4171	4170
293	1781	1780	4173	4172
294	1783	1782	4175	4174
295	1785	1784	4177	4176
296	1787	1786	4179	4178
297	1789	1788	4181	4180
298	1791	1790	4183	4182
299	1793	1792	4185	4184
300	1795	1794	4187	4186
301	1797	1796	4189	4188
302	1799	1798	4191	4190
303	1801	1800	4193	4192
304	1803	1802	4195	4194
305	1805	1804	4197	4196

ORFGenset	ORFoligosFd	ORFoligosFp	ORFaligosBd	ORFoligosBp
306	1807	1806	4199	4198
307	1809	1808	4201	4200
308	1811	1810	4203	4202
309	1813	1812	4205	4204
310	1815	1814	4207	4206
311	1817	1816	4209	4208
312	1819	1818	4211	4210
313	1821	1820	4213	4212
314	1823	1822	4215	4214
315	1825	1824	4217	4216
316	1827	1826	4219	4218
317	1829	1828	4221	4220
318	1831	1830	4223	4222
319	1833	1832	4225	4224
320	1835	1834	4227	4226
321	1837	1836	4229	4228
322	1839	1838	4231	4230
323	1841	1840	4233	4232
324	1843	1842	4235	4234
325	1845	1844	4237	4236
326	1847	1846	4239	4238
327	1849	1848	4241	4240
328	1851	1850	4243	4242
329	1853	1852	4245	4244
330	1855	1854	4247	4246
331	1857	1856	4249	4248
332	1859	1858	4251	4250
333	1861	1860	4253	4252
334	1863	1862	4255	4254
335	1865	1864	4257	4256
336	1867	1866	4259	4258
337	1869	1868	4261	4260
338	1871	1870	426 3	4262
339	1873	1872	4265	4264

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFöligosBp
340	1875	1874	4267	4266
341	1877	1876	4269	4268
342	1879	1878	4271	4270
343	1881	1880	4273	4272
344	1883	1882	4275	4274
345	1885	1884	4277	4276
346	1887	1886	4279	4278
347	1889	1888	4281	4280
348	1891	1890	4283	4282
349	1893	1892	4285	4284
350	1895	1894	4287	4286
351	1897	1896	4289	4288
352	1899	1898	4291	4290
353	1901	1900	4293	4292
354	1903	1902	4295	4294
355	1905	1904	4297	4296
356	1907	1906	4299	4298
357	1909	1908	4301	4300
358	1911	1910	4303	4302
359	1913	1912	4305	4304
360	1915	1914	4307	4306
361	1917	1916	4309	4308
362	1919	1918	4311	4310
363	1921	1920	4313	4312
364	1923	1922	4315	4314
365	1925	1924	4317	4316
366	1927	1926	4319	4318
367	1929	1928	4321	4320
368	1931	1930	4323	4322
369	1933	1932	4325	4324
370	1935	1934	4327	4326
371	1937	1936	4329	4328
372	1939	1938	4331	4330
373	1941	1940	4333	4332

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
374	1943	1942	4335	4334
375	1945	1944	4337	4336
376	1947	1946	4339	4338
377	1949	1948	4341	4340
378	1951	1950	4343	4342
379	1953	1952	4345	4344
380	1955	1954	4347	4346
381	1957	1956	4349	4348
382	1959	1958	4351	4350
383	1961	1960	4353	4352
384	1963	1962	4355	4354
385	1965	1964	4357	4356
386	1967	1966	4359	4358
387	1969	1968	4361	4360
388	1971	1970	4363	4362
389	1973	1972	4365	4364
390	1975	1974	4367	4366
391	1977	1976	4369	4368
392	1979	1978	4371	4370
393	1981	1980	4373	4372
394	1983	1982	4375	4374
395	1985	1984	4377	4376
396	1987	1986	4379	4378
397	1989	1988	4381	4380
398	1991	1990	4383	4382
399	1993	1992	4385	4384
400	1995	1994	4387	4386
401	1997	1996	4389	4388
402	1999	1998	4391	4390
403	2001	2000	4393	4392
404	2003	2002	4395	4394
405	2005	2004	4397	4396
406	2007	2006	4399	4398
407	2009	2008	4401	4400

204					
ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp	
408	2011	2010	4403	4402	
409	2013	2012	4405	4404	
410	2015	2014	4407	4406	
411	2017	2016	4409	4408	
412	2019	2018	4411	4410	
413	2021	2020	4413	4412	
414	2023	2022	4415	4414	
415	2025	2024	4417	4416	
416	2027	2026	4419	4418	
417	2029	2028	4421	4420	
418	2031	2030	4423	4422	
419	2033	2032	4425	4424	
420	2035	2034	4427	4426	
421	2037	2036	4429	4428	
422	2039	2038	4431	4430	
423	2041	2040	4433	4432	
424	2043	2042	4435	4434	
425	2045	2044	4437	4436	
426	2047	2046	4439	4438	
427	2049	2048	4441	4440	
428	2051	2050	4443	4442	
429	2053	2052	4445	4444	
430	2055	2054	4447	4446	
431	2057	2056	4449	4448	
432	2059	2058	4451	4450	
433	2061	2060	4453	4452	
434	2063	2062	4455	4454	
435	2065	2064	4457	4456	
436	2067	2066	4459	4458	
437	2069	2068	4461	4460	
438	2071	2070	4463	4462	
439	2073	2072	4465	4464	
440	2075	2074	4467	4466	
441	2077	2076	4469	4468	
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TANK WELLOW	CONTRACTOR OF THE CONTRACTOR O	ORFGenset ORFoligosEd ORFoligosEn ORFoligosEd ORFoligosEn					
	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp			
442	2079	2078	4471	4470			
443	2081	2080	4473	4472			
444	2083	2082	4475	4474			
445	2085	2084	4477	4476			
446	2087	2086	4479	4478			
447	2089	2088	4481	4480			
448	2091	2090	4483	4482			
449	2093	2092	4485	4484			
450	2095	2094	4487	4486			
451	2097	2096	4489	4488			
452	2099	2098	4491	4490			
453	2101	2100	4493	4492			
454	2103	2102	4495	4494			
455	2105	2104	4497	4496			
456	2107	2106	4499	4498			
457	2109	2108	4501	4500			
458	2111	2110	4503	4502			
459	2113	2112	4505	4504			
460	2115	2114	4507	4506			
461	2117	2116	4509	4508			
462	2119	2118	4511	4510			
463	2121	2120	4513	4512			
464	2123	2122	4515	4514			
465	2125	2124	4517	4516			
466	2127	2126	4519	4518			
467	2129	2128	4521	4520			
468	2131	2130	4523	4522			
469	2133	2132	4525	4524			
470	2135	2134	4527	4526			
471	2137	2136	4529	4528			
472	2139	2138	4531	4530			
473	2141	2140	4533	4532			
474	2143	2142	4535	4534			
475	2145	2144	4537	4536			

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
476	2147	2146	4539	4538
477	2149	2148	4541	4540
478	2151	2150	4543	4542
479	2153	2152	4545	4544
480	2155	2154	4547	4546
481	2157	2156	4549	4548
482	2159	2158	4551	4550
483	2161	2160	4553	4552
484	2163	2162	4555	4554
485	2165	2164	4557	4556
486	2167	2166	4559	4558
487	2169	2168	4561	4560
488	2171	2170	4563	4562
489	2173	2172	4565	4564
490	2175	2174	4567	4566
491	2177	2176	4569	4568
492	2179	2178	4571	4570
493	2181	2180	4573	4572
494	2183	2182	4575	4574
495	2185	2184	4577	4576
496	2187	2186	4579	4578
497	2189	2188	4581	4580
498	2191	2190	4583	4582
499	2193	2192	4585	4584
500	2195	2194	4587	4586
501	2197	2196	4589	4588
502	2199	2198	4591	4590
503	2201	2200	4593	4592
504	2203	2202	4595	4594
505	2205	2204	4597	4596
506	2207	2206	4599	4598
507	2209	2208	4601	4600
508	2211	2210	4603	4602
509	2213	2212	4605	4604

ORFGenset		201		
	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
510	2215	2214	4607	4606
511	2217	2216	4609	4608
512	2219	2218	4611	4610
513	2221	2220	4613	4612
514	2223	2222	4615	4614
515	2225	2224	4617	4616
516	2227	2226	4619	4618
517	2229	2228	4621	4620
518	2231	2230	4623	4622
519	2233	2232	4625	4624
520	2235	2234	4627	4626
521	2237	2236	4629	4628
522	2239	2238	4631	4630
523	2241	2240	4633	4632
524	2243	2242	4635	4634
525	2245	2244	4637	4636
526	2247	2246	4639	4638
527	2249	2248	4641	4640
528	2251	2250	4643	4642
529	2253	2252	4645	4644
530	2255	2254	4647	4646
531	2257	2256	4649	4648
532	2259	2258	4651	4650
533	2261	2260	4653	4652
534	2263	2262	4655	4654
535	2265	2264	4657	4656
536	2267	2266	4659	4658
537	2269	2268	4661	4660
538	2271	2270	4663	4662
539	2273	2272	4665	4664
540	2275	2274	4667	4666
541	2277	2276	4669	4668
542	2279	2278	4671	4670
543	2281	2280	4673	4672
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ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
544	2283	2282	4675	4674
545	2285	2284	4677	4676
546	2287	2286	4679	4678
547	2289	2288	4681	4680
548	2291	2290	4683	4682
549	2293	2292	4685	4684
550	2295	2294	4687	4686
551	2297	2296	4689	4688
552	2299	2298	4691	4690
553	2301	2300	4693	4692
554	2303	2302	4695	4694
555	2305	2304	4697	4696
556	2307	2306	4699	4698
557	2309	2308	4701	4700
558	2311	2310	4703	4702
559	2313	2312	4705	4704
560	2315	2314	4707	4706
561	2317	2316	4709	4708
562	2319	2318	4711	4710
563	2321	2320	4713	4712
564	2323	2322	4715	4714
565	2325	2324	4717	4716
566	2327	2326	4719	4718
567	2329	2328	4721	4720
568	2331	2330	4723	4722
569	2333	2332	4725	4724
570	2335	2334	4727	4726
571	2337	2336	4729	4728
572	2339	2338	4731	4730
573	2341	2340	4733	4732
574	2343	2342	4735	4734
575	2345	2344	4737	4736
576	2347	2346	4739	4738
577	2349	2348	4741	4740

209					
ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp	
578	2351	2350	4743	4742	
579	2353	2352	4745	4744	
580	2355	2354	4747	4746	
581	2357	2356	4749	4748	
582	2359	2358	4751	4750	
583	2361	2360	4753	4752	
584	2363	2362	4755	4754	
585	2365	2364	4757	4756	
586	2367	2366	4759	4758	
587	2369	2368	4761	4760	
588	2371	2370	4763	4762	
589	2373	2372	4765	4764	
590	2375	2374	4767	4766	
591	2377	2376	4769	4768	
592	2379	2378	4771	4770	
593	2381	2380	4773	4772	
594	2383	2382	4775	4774	
595	2385	2384	4777	4776	
596	2387	2386	4779	4778	
597	2389	2388	4781	4780	
598	2391	2390	4783	4782	
599	2393	2392	4785	4784	
600	2395	2394	4787	4786	
601	2397	2396	4789	4788	
602	2399	2398	4791	4790	
603	2401	2400	4793	4792	
604	2403	2402	4795	4794	
605	2405	2404	4797	4796	
606	2407	2406	4799	4798	
607	2409	2408	4801	4800	
608	2411	2410	4803	4802	
609	2413	2412	4805	4804	
610	2415	2414	4807	4806	
611	2417	2416	4809	4808	

ORFGenset	ORFoligösFd	ORFöligösFp	ORFoligosBd	ORFoligosBp
612	2419	2418	4811	4810
613	2421	2420	4813	4812
614	2423	2422	4815	4814
615	2425	2424	4817	4816
616	2427	2426	4819	4818
617	2429	2428	4821	4820
618	2431	2430	4823	4822
619	2433	2432	4825	4824
620	2435	2434	4827	4826
621	2437	2436	4829	4828
622	2439	2438	4831	4830
623	2441	2440	4833	4832
624	2443	2442	4835	4834
625	2445	2444	4837	4836
626	2447	2446	4839	4838
627	2449	2448	4841	4840
628	2451	2450	4843	4842
629	2453	2452	4845	4844
630	2455	2454	4847	4846
631	2457	2456	4849	4848
632	2459	2458	4851	4850
633	2461	2460	4853	4852
634	2463	2462	4855	4854
635	2465	2464	4857	4856
636	2467	2466	4859	4858
637	2469	2468	4861	4860
638	2471	2470	4863	4862
639	2473	2472	4865	4864
640	2475	2474	4867	4866
641	2477	2476	4869	4868
642	2479	2478	4871	4870
643	2481	2480	4873	4872
644	2483	2482	4875	4874
645	2485	2484	4877	4876

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
646	2487	2486	4879	4878
647	2489	2488	4881	4880
648	2491	2490	4883	4882
649	2493	2492	4885	4884
650	2495	2494	4887	4886
651	2497	2496	4889	4888
652	2499	2498	4891	4890
653	2501	2500	4893	4892
654	2503	2502	4895	4894
655	2505	2504	4897	4896
656	2507	2506	4899	4898
657	2509	2508	4901	4900
658	2511	2510	4903	4902
659	2513	2512	4905	4904
660	2515	2514	4907	4906
661	2517	2516	4909	4908
662	2519	2518	4911	4910
663	2521	2520	4913	4912
664	2523	2522	4915	4914
665	2525	2524	4917	4916
666	2527	2526	4919	4918
667	2529	2528	4921	4920
668	2531	2530	4923	4922
669	2533	2532	4925	4924
670	2535	2534	4927	4926
671	2537	2536	4929	4928
672	2539	2538	4931	4930
673	2541	2540	4933	4932
674	2543	2542	4935	4934
675	2545	2544	4937	4936
676	2547	2546	4939	4938
677	2549	2548	4941	4940
678	2551	2550	4943	4942
679	2553	2552	4945	4944

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
680	2555	2554	4947	4946
681	2557	2556	4949	4948
682	2559	2558	4951	4950
683	2561	2560	4953	4952
684	2563	2562	4955	4954
685	2565	2564	4957	4956
686	2567	2566	4959	4958
687	2569	2568	4961	4960
688	2571	2570	4963	4962
689	2573	2572	4965	4964
690	2575	2574	4967	4966
691	2577	2576	4969	4968
692	2579	2578	4971	4970
693	2581	2580	4973	4972
694	2583	2582	4975	4974
695	2585	2584	4977	4976
696	2587	2586	4979	4978
697	2589	2588	4981	4980
698	2591	2590	4983	4982
699	2593	2592	4985	4984
700	2595	2594	4987	4986
701	2597	2596	4989	4988
702	2599	2598	4991	4990
703	2601	2600	4993	4992
704	2603	2602	4995	4994
705	2605	2604	4997	4996
706	2607	2606	4999	4998
707	2609	2608	5001	5000
708	2611	2610	5003	5002
709	2613	2612	5005	5004
710	2615	2614	5007	5006
711	2617	2616	5009	5008
712	2619	2618	5011	5010
713	2621	2620	5013	5012

213					
ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp	
714	2623	2622	5015	5014	
715	2625	2624	5017	5016	
716	2627	2626	5019	5018	
717	2629	2628	5021	5020	
718	2631	2630	5023	5022	
719	2633	2632	5025	5024	
720	2635	2634	5027	5026	
721	2637	2636	5029	5028	
722	2639	2638	5031	5030	
723	2641	2640	5033	5032	
724	2643	2642	5035	5034	
725	2645	2644	5037	5036	
726	2647	2646	5039	5038	
727	2649	2648	5041	5040	
728	2651	2650	5043	5042	
729	2653	2652	5045	5044	
730	2655	2654	5047	5046	
731	2657	2656	5049	5048	
732	2659	2658	5051	5050	
733	2661	2660	5053	5052	
734	2663	2662	5055	5054	
735	2665	2664	5057	5056	
736	2667	2666	5059	5058	
737	2669	2668	5061	5060	
738	2671	2670	5063	5062	
739	2673	2672	5065	5064	
740	2675	2674	5067	5066	
741	2677	2676	5069	5068	
742	2679	2678	5071	5070	
743	2681	2680	5073	5072	
744	2683	2682	5075	5074	
745	2685	2684	5077	5076	
746	2687	2686	5079	5078	
747	2689	2688	5081	5080	

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
748	2691	2690	5083	5082
749	2693	2692	5085	5084
750	2695	2694	5087	5086
751	2697	2696	5089	5088
752	2699	2698	5091	5090
753	2701	2700	5093	5092
754	2703	2702	5095	5094
755	2705	2704	5097	5096
756	2707	2706	5099	5098
757	2709	2708	5101	5100
758	2711	2710	5103	5102
759	2713	2712	5105	5104
760	2715	2714	5107	5106
761	2717	2716	5109	5108
762	2719	2718	5111	5110
763	2721	2720	5113	5112
764	2723	2722	5115	5114
765	2725	2724	5117	5116
766	2727	2726	5119	5118
7 67	2729	2728	5121	5120
768	2731	2730	5123	5122
769	2733	2732	5125	5124
770	2735	2734	5127	5126
771	2737	2736	5129	5128
772	2739	2738	5131	5130
773	2741	2740	5133	5132
774	2743	2742	5135	5134
775	2745	2744	5137	5136
776	2747	2746	5139	5138
777	2749	2748	5141	5140
778	2751	2750	5143	5142
779	2753	2752	5145	5144
780	2755	2754	5147	5146
781	2757	2756	5149	5148

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
782	2759	2758	5151	5150
783	2761	2760	5153	5152
784	2763	2762	5155	5154
785	2765	2764	5157	5156
786	2767	2766	5159	5158
787	2769	2768	5161	5160
788	2771	2770	5163	5162
789	2773	2772	5165	5164
790	2775	2774	5167	5166
791	2777	2776	5169	5168
792	2779	2778	5171	5170
793	2781	2780	5173	5172
794	2783	2782	5175	5174
795	2785	2784	5177	5176
796	2787	2786	5179	5178
79 7	2789	2788	5181	5180
798	2791	2790	5183	5182
799	2793	2792	5185	5184
800	2795	2794	5187	5186
801	2797	2796	5189	5188
802	2799	2798	5191	5190
803	2801	2800	5193	5192
804	2803	2802	5195	5194
805	2805	2804	5197	5196
806	2807	2806	5199	5198
807	2809	2808	5201	5200
808	2811	2810	5203	5202
809	2813	2812	5205	5204
810	2815	2814	5207	5206
811	2817	2816	5209	5208
812	2819	2818	5211	5210
813	2821	2820	5213	5212
814	2823	2822	5215	5214
815	2825	2824	5217	5216

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	Openican
816	2827			ORFoligosBp
		2826	5219	5218
817	2829	2828	5221	5220
818	2831	2830	5223	5222
819	2833	2832	5225	5224
820	2835	2834	5227	5226
821	2837	2836	5229	5228
822	2839	2838	5231	5230
823	2841	2840	5233	5232
824	2843	2842	5235	5234
825	2845	2844	5237	5236
826	2847	2846	5239	5238
827	2849	2848	5241	5240
828	2851	2850	5243	5242
829	2853	2852	5245	5244
830	2855	2854	5247	5246
831	2857	2856	5249	5248
832	2859	2858	5251	5250
833	2861	2860	5253	5252
834	2863	2862	5255	5254
835	2865	2864	5257	5256
836	2867	2866	5259	5258
837	2869	2868	5261	5260
838	2871	2870	5263	5262
839	2873	2872	5265	5264
840	2875	2874	5267	5266
841	2877	2876	5269	5268
842	2879	2878	5271	5270
843	2881	2880	5273	5272
844	2883	2882	5275	5274
845	2885	2884	5277	5276
846	2887	2886	5279	5278
847	2889	2888	5281	5280
848	2891	2890	5283	5282
849	2893	2892	5285	5284
			3203	J20 7

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligesBd	ORFoligosBp
850	2895	2894	5287	5286
851	2897	2896	5289	5288
852	2899	2898	5291	5290
853	2901	2900	5293	5292
854	2903	2902	5295	5294
855	2905	2904	5297	5296
856	2907	2906	5299	5298
857	2909	2908	5301	5300
858	2911	2910	5303	5302
859	2913	2912	5305	5304
860	2915	2914	5307	5306
861	2917	2916	5309	5308
862	2919	2918	5311	5310
863	2921	2920	5313	5312
864	2923	2922	5315	5314
865	2925	2924	5317	5316
866	2927	2926	5319	5318
867	2929	2928	5321	5320
868	2931	2930	5323	5322
869	2933	2932	5325	5324
870	2935	2934	5327	5326
871	2937	2936	5329	5328
872	2939	2938	5331	5330
873	2941	2940	5333	5332
874	2943	2942	5335	5334
875	2945	2944	5337	5336
876	2947	2946	5339	5338
877	2949	2948	5341	5340
878	2951	2950	5343	5342
879	2953	2952	5345	5344
880	2955	2954	5347	5346
881	2957	2956	5349	5348
882	2959	2958	5351	5350
883	2961	2960	5353	5352

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	Openia est
884	2963			ORFoligosBp
885		2962	5355	5354
	2965	2964	5357	5356
886	2967	2966	5359	5358
887	2969	2968	5361	5360
888	2971	2970	5363	5362
889	2973	2972	5365	5364
890	2975	2974	5367	5366
891	2977	2976	5369	5368
892	2979	2978	5371	5370
893	2981	2980	5373	5372
894	2983	2982	5375	5374
895	2985	2984	5377	5376
896	2987	2986	5379	5378
897	2989	2988	5381	5380
898	2991	2990	5383	5382
899	2993	2992	5385	5384
900	2995	2994	5387	5386
901	2997	2996	5389	5388
902	2999	2998	5391	5390
903	3001	3000	5393	5392
904	3003	3002	5395	5394
905	3005	3004	5397	5396
906	3007	3006	5399	5398
907	3009	3008	5401	5400
908	3011	3010	5403	5402
909	3013	3012	5405	5404
910	3015	3012	5407	
911	3017			5406
912	3017	3016	5409	5408
913		3018	5411	5410
913	3021	3020	5413	5412
914	3023	3022	5415	5414
	3025	3024	5417	5416
916	3027	3026	5419	5418
917	3029	3028	5421	5420

ORFGenset	ORFoligesFd	ORFoligosFp	ORFoligosBd	ORFoligosBp		
918	3031	3030	5423	5422		
919	3033	3032	5425	5424		
920	3035	3034	5427	5426		
921	3037	3036	5429	5428		
922	3039	3038	5431	5430		
923	3041	3040	5433	5432		
924	3043	3042	5435	5434		
925	3045	3044	5437	5436		
926	3047	3046	5439	5438		
927	3049	3048	5441	5440		
928	3051	3050	5443	5442		
929	3053	3052	5445	5444		
930	3055	3054	5447	5446		
931	3057	3056	5449	5448		
932	3059	3058	5451	5450		
933	3061	3060	5453	5452		
934	3063	3062	5455	5454		
935	3065	3064	5457	5456		
936	3067	3066	5459	5458		
937	3069	3068	5461	5460		
938	3071	3070	5463	5462		
939	3073	3072	5465	5464		
940	3075	3074	5467	5466		
941	3077	3076	5469	5468		
942	3079	3078	5471	5470		
943	3081	3080	5473	5472		
944	3083	3082	5475	5474		
945	3085	3084	5477	5476		
946	3087	3086	5479	5478		
947	3089	3088	5481	5480		
948	3091	3090	5483	5482		
949	3093	3092	5485	5484		
950	3095	3094	5487	5486		
951	3097	3096	5489	5488		
	 					

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
952	3099	3098	5491	5490
953	3101	3100	5493	5492
954	3103	3102	5495	5494
955	3105	3104	5497	5496
956	3107	3106	5499	5498
957	3109	3108	5501	5500
958	3111	3110	5503	5502
959	3113	3112	5505	5504
960	3115	3114	5507	5506
961	3117	3116	5509	5508
962	3119	3118	5511	5510
963	3121	3120	5513	5512
964	3123	3122	5515	5514
965	3125	3124	5517	5516
966	3127	3126	5519	5518
967	3129	3128	5521	5520
968	3131	3130	5523	5522
969	3133	3132	5525	5524
970	3135	3134	5527	5526
971	3137	3136	5529	5528
972	3139	3138	5531	5530
973	3141	3140	5533	5532
974	3143	3142	5535	5534
975	3145	3144	5537	5536
976	3147	3146	5539	5538
977	3149	3148	5541	5540
978	3151	3150	5543	5542
979	3153	3152	5545	5544
980	3155	3154	5547	5546
981	3157	3156	5549	5548
982	3159	3158	5551	5550
983	3161	3160	5553	5552
984	3163	3162	5555	5554
985	3165	3164	5557	5556

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ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp		
986	3167	3166	5559	5558		
987	3169	3168	5561	5560		
988	3171	3170	5563	5562		
989	3173	3172	5565	5564		
990	3175	3174	5567	5566		
991	3177	3176	5569	5568		
992	3179	3178	5571	5570		
993	3181	3180	5573	5572		
994	3183	3182	5575	5574		
995	3185	3184	5577	5576		
996	3187	3186	5579	5578		
997	3189	3188	5581	5580		
998	3191	3190	5583	5582		
999	3193	3192	5585	5584		
1000	3195	3194	5587	5586		
1001	3197	3196	5589	5588		
1002	3199	3198	5591	5590		
1003	3201	3200	5593	5592		
1004	3203	3202	5595	5594		
1005	3205	3204	5597	5596		
1006	3207	3206	5599	5598		
1007	3209	3208	5601	5600		
1008	3211	3210	5603	5602		
1009	3213	3212	5605	5604		
1010	3215	3214	5607	5606		
1011	3217	3216	5609	5608		
1012	3219	3218	5611	5610		
1013	3221	3220	5613	5612		
1014	3223	3222	5615	5614		
1015	3225	3224	5617	5616		
1016	3227	3226	5619	5618		
1017	3229	3228	5621	5620		
1018	3231	3230	5623	5622		
1019	3233	3232	5625	5624		

ORFGenset	ORFoligosFd	ADE-HASE		
		ORFoligosFp	ORFoligosBd	ORFoligosBp
1020	3235	3234	5627	5626
1021	3237	3236	5629	5628
1022	3239	3238	5631	5630
1023	3241	3240	5633	5632
1024	3243	3242	5635	5634
1025	3245	3244	5637	5636
1026	3247	3246	5639	5638
1027	3249	3248	5641	5640
1028	3251	3250	5643	5642
1029	3253	3252	5645	5644
1030	3255	3254	5647	5646
1031	3257	325ύ	5649	5648
1032	3259	3258	5651	5650
1033	3261	3260	5653	5652
1034	3263	3262	5655	5654
1035	3265	3264	5657	5656
1036	3267	3266	5659	5658
1037	3269	3268	5661	5660
1038	3271	3270	5663	5662
1039	3273	3272	5665	5664
1040	3275	3274	5667	5666
1041	3277	3276	5669	5668
1042	3279	3278	5671	5670
1043	3281	3280	5673	5672
1044	3283	3282	5675	5674
1045	3285	3284	5677	5676
1046	3287	3286	5679	5678
1047	3289	3288	5681	5680
1048	3291	3290	5683	5682
1049	3293	3292	5685	5684
1050	3295	3294	5687	5686
1051	3297	3296	5689	5688
1052	3299	3298	5691	5690
1053	3301	3300	5693	5692

223					
ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligasBp	
1054	3303	3302	5695	5694	
1055	3305	3304	5697	5696	
1056	3307	3306	5699	5698	
1057	3309	3308	5701	5700	
1058	3311	3310	5703	5702	
1059	3313	3312	5705	5704	
1060	3315	3314	5707	5706	
1061	3317	3316	5709	5708	
1062	3319	3318	5711	5710	
1063	3321	3320	5713	5712	
1064	3323	3322	5715	5714	
1065	3325	3324	5717	5716	
1066	3327	3326	5719	5718	
1067	3329	3328	5721	5720	
1068	3331	3330	5723	5722	
1069	3333	3332	5725	5724	
1070	3335	3334	5727	5726	
1071	3337	3336	5729	5728	
1072	3339	3338	5731	5730	
1073	3341	3340	5733	5732	
1074	3343	3342	5735	5734	
1075	3345	3344	5737	5736	
1076	3347	3346	5739	5738	
1077	3349	3348	5741	5740	
1078	3351	3350	5743	5742	
1079	3353	3352	5745	5744	
1080	3355	3354	5747	5746	
1081	3357	3356	5749	5748	
1082	3359	3358	5751	5750	
1083	3361	3360	5753	5752	
1084	3363	3362	5755	5754	
1085	3365	3364	5757	5756	
1086	3367	3366	5759	5758	
1087	3369	3368	5761	5760	

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1088	3371	3370	5763	5762
1089	3373	3372	5765	5764
1090	3375	3374	5767	5766
1091	3377	3376	5769	5768
1092	3379	3378	5771	5770
1093	3381	3380	5773	5772
1094	3383	3382	5775	5774
1095	3385	3384	5777	5776
1096	3387	3386	5779	5778
1097	3389	3388	5781	5780
1098	3391	3390	5783	5782
1099	3393	3392	5785	5784
1100	3395	3394	5787	5786
1101	3397	3396	5789	5788
1102	3399	3398	5791	5790
1103	3401	3400	5793	5792
1104	3403	3402	5795	5794
1105	3405	3404	5797	5796
1106	3407	3406	5799	5798
1107	3409	3408	5801	5800
1108	3411	3410	5803	5802
1109	3413	3412	5805	5804
1110	3415	3414	5807	5806
1111	3417	3416	5809	5808
1112	3419	3418	5811	5810
1113	3421	3420	5813	5812
1114	3423	3422	5815	5814
1115	3425	3424	5817	5816
1116	3427	3426	5819	5818
1117	3429	3428	5821	5820
1118	3431	3430	5823	5822
1119	3433	3432	5825	5824
1120	3435	3434	5827	5826
1121	3437	3436	5829	5828

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1122	3439	3438	5831	5830
1123	3441	3440	5833	5832
1124	3443	3442	5835	5834
1125	3445	3444	5837	5836
1126	3447	3446	5839	5838
1127	3449	3448	5841	5840
1128	3451	3450	5843	5842
1129	3453	3452	5845	5844
1130	3455	3454	5847	5846
1131	3457	3456	5849	5848
1132	3459	3458	5851	5850
1133	3461	3460	5853	5852
1134	3463	3462	5855	5854
1135	3465	3464	5857	5856
1136	3467	3466	5859	5858
1137	3469	3468	5861	5860
1138	3471	3470	5863	5862
1139	3473	3472	5865	5864
1140	3475	3474	5867	5866
1141	3477	3476	5869	5868
1142	3479	3478	5871	5870
1143	3481	3480	5873	5872
1144	3483	3482	5875	5874
1145	3485	3484	5877	5876
1146	3487	3486	5879	5878
1147	3489	3488	5881	5880
1148	3491	3490	5883	5882
1149	3493	3492	5885	5884
1150	3495	3494	5887	5886
1151	3497	3496	5889	5888
1152	3499	3498	5891	5890
1153	3501	3500	5893	5892
1154	3503	3502	5895	5894
1155	3505	3504	5897	5896

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ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp	
1156	3507	3506	5899	5898	
1157	3509	3508	5901	5900	
1158	3511	3510	5903	5902	
1159	3513	3512	5905	5904	
1160	3515	3514	5907	5906	
1161	3517	3516	5909	5908	
1162	3519	3518	5911	5910	
1163	3521	3520	5913	5912	
1164	3523	3522	5915	5914	
1165	3525	3524	5917	5916	
1166	3527	3526	5919	5918	
1167	3529	3528	5921	5920	
1168	3531	3530	5923	5922	
1169	3533	3532	5925	5924	
1170	3535	3534	5927	5926	
1171	3537	3536	5929	5928	
1172	3539	3538	5931	5930	
1173	3541	3540	5933	5932	
1174	3543	3542	5935	5934	
1175	3545	3544	5937	5936	
1176	3547	3546	5939	5938	
1177	3549	3548	5941	5940	
1178	3551	3550	5943	5942	
1179	3553	3552	5945	5944	
1180	3555	3554	5947	5946	
1181	3557	3556	5949	5948	
1182	3559	3558	5951	5950	
1183	3561	3560	5953	5952	
1184	3563	3562	5955	5954	
1185	3565	3564	5957	5956	
1186	3567	3566	5959	5958	
1187	3569	3568	5961	5960	
1188	3571	3570	5963	5962	
1189	3573	3572	5965	5964	

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1190	3575	3574	5967	5966
1191	3577	3576	5969	5968
1192	3579	3578	5971	5970
1193	3581	3580	5973	5972
1194	3583	3582	5975	5974
1195	3585	3584	5977	5976
1196	3587	3586	5979	5978
1197	3589	3588	5981	5980

TABLE 5

SEQ ID	Or.	position
1198	F	1038449
1199	F	1036517
1200	F	250
1201	F	1036965
1202	F	3011
1203	F	1123
1204	F	4907
1205	F	2996
1206	F	6379
1207	F	4483
1208	F	7837
1209	F	5961
1210	F	8351
1211	F	6467
1212	F	8705
1213	F	6834
1214	F	9598
1215	F	7709
1216	F	10134
1217	F	8248
1218	F	10990
1219	F	9060
1220	F	11823
1221	F	9946
1222	F	13236
1223	F	11410
1224	F	14529
1225	F	12643
1226	F	14668
1227	F	12813
1228	F	15747
1229	F	13844

SEQ ID	Or.	position		
2793	F	785793		
2794	F	789918		
2795	F	788039		
2796	F	790378		
2797	F	788456		
2798	F	791834		
2799	F	789918		
2800	F	793102		
2801	F	791176		
2802	F	793826		
2803	F	791921		
2804	F	794911		
2805	F	793023		
2806	F	795296		
2807	F	793427		
2808	F	796005		
2809	F	794127		
2810	F	796729		
2811	F	794811		
2812	F	797041		
2813	F	795065		
2814	F	797553		
2815	F	795651		
2816	Ŧ	797716		
2817	F	795815		
2818	F	798197		
2819	F	796285		
2820	F	799004		
2821	F	797173		
2822	F	799785		
2823	F	797910		
2824	F	800789		

4388 B 394245 4389 B 396116 4390 B 395604 4391 B 397475 4392 B 396249 4393 B 398133 4394 B 396759 4395 B 398660 4396 B 397746 4397 B 399639 4398 B 398973 4399 B 400878 4400 B 399921 4401 B 401846 4402 B 400393 4403 B 402287 4404 B 401444 4405 B 403344 4406 B 402258 4407 B 404150 4408 B 403461 4409 B 405400 4411 B 406141 4412 B 40627 4413 B 405941 4414 B 406141 <tr< th=""><th>SEQ ID</th><th>Or.</th><th>position</th></tr<>	SEQ ID	Or.	position
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230				
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	1432	F	119580
	1433	F	117660

234			
SEQ ID	Or.	position	
2995	F	876963	
2996	F	879824	
2997	F	877933	
2998	F	880670	
2999	F	878769	
3000	F	881719	
3001	F	879824	
3002	F	882682	
3003	F	880774	
3004	F	883432	
3005	F	881540	
3006	F	884263	
3007	F	882357	
3008	F	884947	
3009	F	883044	
3010	F	888721	
3011	F	886762	
3012	F	890084	
3013	F	888182	
3014	F	890897	
3015	F	888996	
3016	F	891749	
3017	F	889830	
3018	F	893136	
3019	F	891228	
3020	F	893415	
3021	F	891471	
3022	F	893591	
3023	F	891684	
3024	F	894005	
3025	F	892127	
3026	F	894827	
3027	F	892900	
3028	F	895732	

SEQ ID	Or.	position
4590	В	496854
4591	В	498754
4592	В	497396
4593	В	499316
4594	В	498735
4595	В	500635
4596	В	499484
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4598	В	501005
4599	В	502852
4600	В	501937
4601	В	503853
4602	В	503083
4603	В	505003
4604	В	503895
4605	В	505846
4606	В	505263
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4608	В	507214
4609	В	509106
4610	В	507687
4611	В	509559
4612	В	508632
4613	В	510534
4614	В	508863
4615	В	510730
4616	В	509202
4617	В	511062
4618	В	510940
4619	В	512832
4620	В	511747
4621	В	513649
4622	В	512446
4623	В	514305

SEQ ID	Or.	position
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1436	F	124649
1437	F	122753
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1439	F	123416
1440	F	126101
1441	F	124208
1442	F	126871
1443	F	125013
1444	F	127698
1445	F	125787
1446	F	129465
1447	F	127467
1448	F	130799
1449	F	128869
1450	F	131615
1451	F	129711
1452	F	132856
1453	F	130914
1454	F	133401
1455	F	131474
1456	F	133624
1457	F	131706
1458	F	134385
1459	F	132500
1460	F	137183
1461	F	135320
1462	F	140106
1463	F	138215
1464	F	140839
1465	F	138927
1466	F	141535
1467	F	139614

235			
SEQ ID	Or.	position	
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3030	F	896823	
3031	F	894907	
3032	F	900571	
3033	F	898639	
3034	F	902407	
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3036	F	903243	
3037	F	901346	
3038	F	903616	
3039	F	901726	
3040	F	905486	
3041	F	903589	
3042	F	906234	
3043	F	904350	
3044	F	906774	
3045	F	904846	
3046	F	907868	
3047	F	905924	
3048	F	908501	
3049	F	906583	
3050	F	908975	
3051	F	907079	
3052	F	909351	
3053	F	907456	
3054	F	909835	
3055	F	907957	
3056	F	910382	
3057	F	908496	
3058	F	910693	
3059	F	908829	
3060	F	912169	
3061	F	910248	
3062	F	912376	

SEQ ID	10	l
	Or.	
4624	В	513313
4625	В	515212
4626	В	514529
4627	В	516410
4628	В	515466
4629	В	517364
4630	В	515496
4631	В	517389
4632	В	516069
4633	В	517978
4634	В	516642
4635	В	518551
4636	В	517420
4637	В	519349
4638	В	518187
4639	В	520053
4640	В	518617
4641	В	520500
4642	В	519078
4643	В	520963
4644	В	519736
4645	В	521636
4646	В	520719
4647	В	522655
4648	В	522221
4649	В	524115
4650	В	522354
4651	В	524287
4652	В	523763
4653	В	525689
4654	В	524854
4655	В	526756
4656	В	525970
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SEQ ID	Or.	position
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1470	F	143684
1471	F	141782
1472	F	144309
1473	F	142375
1474	F	146178
1475	F	144294
1476	F	146894
1477	F	144997
1478	F	147858
1479	F	145960
1480	F	148277
1481	F	146347
1482	F	148781
1483	F	146846
1484	F	148947
1485	F	147021
1486	F	149424
1487	F	147592
1488	F	150769
1489	F	148884
1490	F	151743
1491	F	149880
1492	F	152659
1493	F	150769
1494	F	153101
1495	F	151270
1496	F	153719
1497	F	151850
1498	F	155002
1499	F	153096
1500	F	156550
1501	F	154687

SEQ ID	Or.	position
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3065	F	911084
3066	F	913437
3067	F	911545
3068	F	914282
3069	F	912376
3070	F	914925
3071	F	913023
3072	F	915394
3073	F	913510
3074	F	915827
3075	F	913912
3076	F	916683
3077	F	914788
3078	F	917347
3079	F	915438
3080	F	918089
3081	F	916189
3082	F	918399
3083	F	916506
3084	F	919296
3085	F	917406
3086	F	919457
3087	F	917598
3088	F	919864
3089	F	917963
3090	F	920641
3091	F	918711
3092	F	921029
3093	F	919138
3094	F	921239
3095	F	919366
3096	F	921526

SEQ ID		
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4660	В	526640
4661	В	528553
4662	В	526991
4663	В	528855
4664	В	528553
4665	В	530443
4666	В	529081
4667	В	530988
4668	В	529943
4669	В	531844
4670	В	530424
4671	В	532301
4672	В	530799
4673	В	532675
4674	В	531670
4675	В	533594
4676	В	533498
4677	В	535393
4678	В	534147
4679	В	535997
4680	В	534892
4681	В	536813
4682	В	536191
4683	В	538068
4684	В	539438
4685	В	541306
4686	В	540771
4687	В	542639
4688	В	541223
4689	В	543141
4690	В	542025
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SEQ ID	Or.	position
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1503	F	155353
1504	F	158818
1505	F	156924
1506	F	159676
1507	F	157795
1508	F	160957
1509	F	159063
1510	F	161319
1511	F	159504
1512	F	162131
1513	F	160240
1514	F	162775
1515	F	160865
1516	F	164236
1517	F	162345
1518	F	165837
1519	F	163923
1520	F	166508
1521	F	164605
1522	F	168612
1523	F	166683
1524	F	169367
1525	F	167436
1526	F	170556
1527	F	168661
1528	F	171067
1529	F	169173
1530	F	172090
1531	F	170171
1532	F	172797
1533	F	170904
1534	F	174176
1535	F	172236

201			
SEQ ID	Or.	position	
3097	F	919638	
3098	F	921930	
3099	F	919979	
3100	F	922212	
3101	F	920325	
3102	F	922925	
3103	F	921029	
3104	F	923258	
3105	F	921324	
3106	F	923808	
3107	F	921929	
3108	F	924185	
3109	F	922311	
3110	F	924680	
3111	F	922764	
3112	F	925111	
3113	F	923258	
3114	F	926538	
3115	F	924638	
3116	F	926972	
3117	F	925072	
3118	F	927351	
3119	F	925415	
3120	F	927870	
3121	F	925924	
3122	F	928974	
3123	F	927031	
3124	F	930003	
3125	F	928103	
3126	F	930383	
3127	F	928402	
3128	F	931084	
3129	F	929222	
3130	F	931307	

SEQ ID	10-	langi ti aa
	Or.	
4692	В	543495
4693	В	545375
4694	В	544367
4695	В	546253
4696	В	544790
4697	В	546697
4698	В	544982
4699	В	546890
4700	В	546655
4701	В	548555
4702	В	547701
4703	В	549667
4704	В	547609
4705	В	549533
4706	В	548121
4707	В	550040
4708	В	548878
4709	В	550836
4710	В	549681
4711	В	551602
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4713	В	552527
4714	В	551849
4715	В	553750
4716	В	553261
4717	В	555165
4718	В	555176
4719	В	557075
4720	В	556590
4721	В	558489
4722	В	557130
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SEQ ID	Or.	position
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1538	F	175476
1539	F	173595
1540	F	177183
1541	F	175275
1542	F	177858
1543	F	175942
1544	F	179819
1545	F	177923
1546	F	180412
1547	F	178455
1548	F	181169
1549	F	179226
1550	F	182628
1551	F	180694
1552	F	183403
1553	F	181494
1554	F	184577
1555	F	182628
1556	F	185763
1557	F	183843
1558	F	186496
1559	F	184614
1560	F	187187
1561	F	185262
1562	F	188542
1563	F	186632
1564	F	189410
1565	F	187514
1566	F	190016
1567	F	188083
1568	F	190545
1569	F	188666

238			
SEQ ID	Or.	position	
3131	F	929397	
3132	F	931824	
3133	F	929927	
3134	F	932352	
3135	F	930470	
3136	F	933044	
3137	F	931084	
3138	F	933303	
3139	F	931396	
3140	F	933626	
3141	F	931686	
3142	F	934320	
3143	F	932412	
3144	F	936427	
3145	F	934508	
3146	F	938309	
3147	F	936402	
3148	F	939110	
3149	F	937204	
3150	F	940791	
3151	F	938889	
3152	F	941806	
3153	F	939906	
3154	F	944314	
3155	F	942412	
3156	F	944987	
3157	F	943090	
3158	F	946072	
3159	F	944166	
3160	F	946877	
3161	F	944986	
3162	F	948258	
3163	F	946403	
3164	F	949037	

SEQ ID	Or.	position
4726	В	558455
4727	В	560376
4728	В	559450
4729	В	561362
4730	В	561581
4731	В	563478
4732	В	563153
4733	В	565073
4734	В	564319
4735	В	566220
4736	В	564201
4737	В	566078
4738	В	567243
4739	В	569143
4740	В	568192
4741	В	570133
4742	В	570619
4743	В	572532
4744	В	572241
4745	В	574208
4746	В	572994
4747	В	574916
4748	В	573744
4749	В	575640
4750	В	573679
4751	В	575571
4752	В	574398
4753	В	576288
4754	В	574677
4755	В	576633
4756	В	575033
4757	В	576922
4758	В	575482
4759	В	577363

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[SEQ ID	Or.	position	
	1570	F	191538	
	1571	F	189595	
	1572	F	192173	
•	1573	F	190247	
	1574	F	193015	
ļ	1575	F	191135	
Ī	1576	F	194471	
	1577	F	192522	
	1578	F	194946	
Ī	1579	F	193015	
Ī	1580	F	196798	
	1581	F	194896	
Ī	1582	F	197440	
	1583	F	195550	
	1584	F	197440	
ſ	1585	F	195549	
	1586	F	198736	
	1587	F	196802	
Ī	1588	F	199722	
Ī	1589	F	197822	
ſ	1590	F	200003	
	1591	F	198147	
	1592	F	200361	
	1593	F	198453	
	1594	F	200945	
	1595	F	199009	
Γ	1596	F	202122	
	1597	F	200215	
	1598	F	203251	
	1599	F	201352	
Γ	1600	F	203807	
	1601	F	201903	
	1602	F	206555	
	1603	F	204669	

	239			
SEQ		Or.	position	
316		F	947137	
316	6	F	949581	
316	7	F	947671	
316	8	F	950455	
316	9	F	948558	
317	0	F	951058	
317	1	F	949151	
317	2	F	951569	
317	3	F	949653	
317	4	F	953340	
317	5	F	951431	
317	6	F	954174	
317	7	F	952288	
317	8	F	955475	
317	9	F	953604	
318	0	F	957601	
318	1	F	955703	
318	2	F	959053	
318	3	F	957168	
318	4	F	960824	
318	5	F	958932	
318	6	F	961268	
318	7	F	959368	
318	8	F	961365	
318	9	F	959465	
319	0	F	962324	
319	1	F	960423	
319	2	F	964706	
319	3	F	962822	
319	4	F	965974	
319	5	F	964074	
319	5	F	967210	
319	7	F	965307	
319	3	F	967808	
				

		
SEQ ID	Or.	position
4760	В	575699
4761	В	577606
4762	В	576078
4763	В	577993
4764	В	578265
4765	В	580143
4766	В	578948
4767	В	580848
4768	В	582336
4769	В	584225
4770	В	582917
4771	В	584317
4772	В	583359
4773	В	585252
4774	В	583734
4775	В	585634
4776	В	584122
4777	В	585990
4778	В	584665
4779	В	586598
4780	В	585213
4781	В	587123
4782	В	585517
4783	В	587434
4784	В	586850
4785	В	588730
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4789	В	591543
4790	В	589798
4791	В	591723
4792	В	590323
4793	В	592211

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	SEQ ID	Or.	position
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	1605	F	205369
	1606	F	208293
	1607	F	206395
	1608	F	209252
•	1609	F	207345
	1610	F	210330
	1611	F	208414
	1612	F	210632
	1613	F	208694
	1614	F	211151
	1615	F	209255
	1616	F	212650
	1617	F	210756
	1618	F	213920
	1619	F	212036
	1620	F	214535
	1621	F	212635
	1622	F	215003
	1623	F	213077
	1624	F	216641
	1625	F	214772
	1626	F	216869
	1627	F	214961
	1628	F	218145
	1629	F	216218
	1630	F	218461
	1631	F	216576
	1632	F	218960
	1633	F	217044
	1634	F	219646
Ì	1635	F	217772
ľ	1636	F	220257
	1637	F	218379

SEQ ID	Or.	position		
3199	F	965925		
3200	F	969286		
3201	F	967320		
3202	F	970802		
3203	F	968888		
3204	F	972169		
3205	F	970269		
3206	F	973487		
3207	F	971616		
3208	F	974339		
3209	F	972408		
3210	F	974988		
3211	F	973035		
3212	F	976035		
3213	F	974114		
3214	F	976367		
3215	F	974411		
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3217	F	974730		
3218	F	977439		
3219	F	975500		
3220	F	977698		
3221	F	975799		
3222	F	978389		
3223	F	976478		
3224	F	978665		
3225	F	976760		
3226	F	979134		
3227	F	977270		
3228	F	979473		
3229	F	977592		
3230	F	980915		
3231	F	979026		
3232	F	982043		

1501		position
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4795	В	593419
4796	В	593002
4797	В	594885
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4806	В	597147
4807	В	598998
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4809	В	599851
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4811	В	602929
4812	В	602096
4813	В	603996
4814	В	603761
4815	В	605643
4816	В	604014
4817	В	605920
4818	В	604634
4819	В	606548
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4822	В	606903
4823	В	608742
4824	В	607722
4825	В	609674
4826	В	609329
4827	В	611215

SEQ ID	Or.	position
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1639	F	218989
1640	F	221314
1641	F	219470
1642	F	222253
1643	F	220338
1644	F	223186
1645	F	221278
1646	F	223994
1647	F	222146
1648	F	224908
1649	F	223014
1650	F	225051
1651	F	223131
1652	F	225510
1653	F	223615
1654	F	226550
1655	F	224609
1656	F	226928
1657	F	225029
1658	F	227528
1659	F	225631
1660	F	228388
1661	F	226475
1662	F	229930
1663	F	228032
1664	F	231129
1665	F	229205
1666	F	232785
1667	F	230915
1668	F	233561
1669	F	231664
1670	F	234013
1671	F	232149

	471	
SEQ ID	Or.	position
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3234	F	983628
3235	F	981736
3236	F	984149
3237	F	982271
3238	F	985180
3239	F	983280
3240	F	985815
3241	F	983882
3242	F	986458
3243	F	984547
3244	F	987340
3245	F	985462
3246	F	987686
3247	F	985815
3248	F	988559
3249	F	986670
3250	F	989455
3251	F	987558
3252	F	993122
3253	F	991289
3254	F	993186
3255	F	991297
3256	F	993650
3257	F	991727
3258	F	994007
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3260	F	995743
3261	F	993915
3262	F	996663
3263	F	994763
3264	F	998586
3265	F	996755
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4882 B 638311 4883 B 640204	4880	В	632046
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	4882	В	638311
4884 B 640448	4883	В	640204
	4884	В	640448
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4887 B 645639	4887	В	645639
4888 B 640847	4888	В	640847
4889 B 642769	4889	В	642769
4890 B 644745	4890	В	644745
4891 B 646615	4891	В	646615
4892 B 645686	4892	В	645686
4893 B 647558	4893	В	647558
4894 B 646060	4894	В	646060
4895 B 647972	4895	В	647972

SEQ ID	Or.	position
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1708	F	257692
1709	F	255790
1710	F	258561
1711	F	256651
1712	F	258927
1713	F	257036
1714	F	261368
1715	F	259469
1716	F	263887
1717	F	262000
1718	F	264503
1719	F	262599
1720	F	265364
1721	F	263512
1722	F	266202
1723	F	264277
1724	F	266709
1725	F	264801
1726	F	267847
1727	F	265947
1728	F	267980
1729	F	266077
1730	F	268271
1731	F	266341
1732	F	269840
1733	F	267913
1734	F	270961
1735	F	269072
1736	F	271883
1737	F	270080
1738	F	272642
1739	F	270748

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SEQ ID	Or.	position
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3303	F	1013010
3304	F	1015664
3305	F	1013820
3306	F	1017026
3307	F	1015099
3308	F	1017674
3309	F	1015786
3310	F	1018353
3311	F	1016460
3312	F	1019602
3313	F	1017674
3314	F	1019876
3315	F	1017948
3316	F	1020853
3317	F	1018956
3318	F	1021878
3319	F	1019972
3320	F	1023054
3321	F	1021186
3322	F	1023415
3323	F	1021579
3324	F	1023748
3325	F	1021850
3326	F	1024485
3327	F	1022574
3328	F	1024744
3329	F	1022836
3330	F	1025618
3331	F	1023720
3332	F	1026323
3333	F	1024403
3334	F	1027710

SEQ ID	Or.	position
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4898		649987
4899	B	651829
4900	В	650580
4901	B	652484
4902	В	651942
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4904	В	652395
4905	В	654310
4906	В	653132
4907	В	655028
4908	В	653827
4909	В	655713
4910	В	662071
4911	В	664023
4912	В	662543
4913	В	664403
4914	В	663295
4915	В	665205
4916	В	663972
4917	В	665850
4918	В	664432
4919	В	666332
4920	В	665860
4921	В	667789
4922	В	666312
4923	В	668233
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4929	В	670495
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SEQ ID	Or.	position
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1741	F	271477
1742	F	274562
1743	F	272702
1744	F	275882
1745	F	273984
1746	F	278004
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1749	F	276893
1750	F	279521
1751	F	277632
1752	F	281076
1753	F	279118
1754	F	281551
1755	F	279668
1756	F	282573
1757	F	280663
1758	F	284229
1759	F	282316
1760	F	284598
1761	F	282655
1762	F	285418
1763	F	283518
1764	F	286104
1765	F	284229
1766	F	286456
1767	F	284531
1768	F	287865
1769	F	286008
1770	F	289163
1771	F	287384
1772	F	290609
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SEQ ID	Or.	position
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3336	F	1030272
3337	F	1028389
3338	F	1031486
3339	F	1029602
3340	F	1033215
3341	F	1031334
3342	F	1035425
3343	F	1033555
3344	F	1035956
3345	F	1034055
3346	F	1036748
3347	F	1034844
3348	F	16372
3349	F	14463
3350	F	31184
3351	F	29287
3352	F	56283
3353	F	54383
3354	F	56384
3355	F	54538
3356	F	64528
3357	F	62600
3358	F	72965
3359	F	71054
3360	F	78245
3361	F	76347
3362	F	79133
3363	F	77291
3364	F	81740
3365	F	79840
3366	F	86772
3367	F	84880
3368	F	109188

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SEQ ID	Or.	position
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4931	В	670590
4932	В	669766
4933	В	671653
4934	В	670160
4935	В	672109
4936	В	671000
4937	В	672900
4938	В	671470
4939	В	673412
4940	В	672685
4941	В	674567
4942	В	673461
4943	В	675365
4944	В	674786
4945	В	676682
4946	В	675456
4947	В	677375
4948	В	676683
4949	В	678594
4950	В	677334
4951	В	679183
4952	В	678726
4953	В	680596
4954	В	679729
4955	В	681628
4956	В	680747
4957	В	682668
4958	В	681500
4959	В	683406
4960	В	682779
4961	В	684716
4962	В	683320
4963	В	685249

SEQ ID	Or.	position
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1776	F	292107
1777	F	290166
1778	F	293099
1779	F	291211
1780	F	294791
1781	F	292883
1782	F	295464
1783	F	293573
1784	F	296018
1785	F	294095
1786	F	297572
1787	F	295664
1788	F	298686
1789	F	296716
1790	F	300305
1791	F	298407
1792	F	301852
1793	F	299946
1794	F	304754
1795	F	302849
1796	F	305854
1797	F	303992
1798	F	306214
1799	F	304303
1800	F	306758
1801	F	304856
1802	F	309057
1803	·F	307125
1804	F	309635
1805	F	307750
1806	F	310491
1807	F	308597

SEQ ID	Or.	position
3369	F	107337
3370	F	111132
3371	F	109188
3372	F	111505
3373	F	109597
3374	F	112432
3375	F	110462
3376	F	113446
3377	F	111592
3378	F	120225
3379	F	118303
3380	F	124892
3381	F	123004
3382	F	131327
3383	F	129485
3384	F	143944
3385	F	142043
3386	F	150138
3387	F	148247
3388	F	163715
3389	F	161804
3390	F	165186
3391	F	163274
3392	F	168143
3393	F	166302
3394	F	170287
3395	F	168387
3396	F	176838
3397	F	174996
3398	F	187776
3399	F	185900
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3402	F	190117

SEQ ID	Or.	position
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4972	В	688619
4973	В	690482
4974	В	688653
4975	В	690563
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4979	В	692088
4980	В	690715
4981	В	692616
4982	В	690937
4983	В	692837
4984	В	692091
4985	В	693991
4986	В	694171
4987	В	696078
4988	В	695197
4989	В	697093
4990	В	697486
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SEQ ID	Or.	position
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1810	F	313188
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1812	F	314121
1813	F	312194
1814	F	314489
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1816	F	315431
1817	F	313526
1818	F	316309
1819	F	314380
1820	F	317102
1821	F	315214
1822	F	317271
1823	F	315343
1824	F	317380
1825	F	315480
1826	F	318256
1827	F	316352
1828	F	319047
1829	F	317181
1830	F	320325
1831	F	318338
1832	F	321228
1833	F	319366
1834	F	321676
1835	F	319782
1836	F	322066
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1838	F	322910
1839	F	320982
1840	F	324744
1841	F	322849

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SEQ ID	Or.	position
3403	F	188168
3404	F	196802
3405	F	194946
3406	F	210685
3407	F	208785
3408	F	234633
3409	F	232727
3410	F	236682
3411	F	234794
3412	F	249227
3413	F	247310
3414	F	252939
3415	F	251036
3416	F	253406
3417	F	251562
3418	F	271365
3419	F	269466
3420	F	275390
3421	F	273489
3422	F	277681
3423	F	275765
3424	F	282260
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3426	F	292925
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3428	F	302910
3429	F	301032
3430	F	308746
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3432	F	311994
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3434	F	312375
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SEQ ID	Or.	position
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4999	В	703015
5000	В	702385
5001	В	704285
5002	В	703636
5003	В	705561
5004	В	705271
5005	В	707136
5006	В	705875
5007	В	707725
5008	В	706444
5009	В	708279
5010	В	706741
5011	В	708673
5012	В	708324
5013	В	710226
5014	В	708673
5015	В	710518
5016	В	708876
5017	В	710791
5018	В	710498
5019	В	712447
5020	В	711435
5021	В	713354
5022	В	712993
5023	В	714887
5024	В	713686
5025	В	715574
5026	В	714474
5027		716354
5028	В	714867
5029	В	716760
5030	В	716047
5031	В	717877

SEQ ID	Or.	position
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1843	F	323445
1844	F	326217
1845	F	324331
1846	F	327038
1847	F	325162
1848	F	327957
1849	F	326079
1850	F	328458
1851	F	326612
1852	F	329032
1853	F	327173
1854	F	329329
1855	F	327489
1856	F	330446
1857	F	328551
1858	F	330915
1859	F	329032
1860	F	331410
1861	F	329602
1862	F	332534
1863	F	330626
1864	F	332782
1865	F	330879
1866	F	333587
1867	F	331632
1868	F	333870
1869	F	331962
1870	F	334510
1871	F	332594
1872	F	334958
1873	F	333049
1874	F	334958
1875	F	333049

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SEQ ID	Or.	
3437	F	310647
3438	F	319923
3439	F	318009
3440	F	339991
3441	F	338104
3442	F	352535
3443	F	350653
3444	F	373218
3445	F	371320
3446	F	376994
3447	F	375085
3448	F	378954
3449	F	377011
3450	F	394604
3451	F	392704
3452	F	400915
3453	F	398972
3454	F	409744
3455	F	407904
3456	F	411155
3457	F	409253
3458	F	414197
3459	F	412281
3460	F	422638
3461	F	420770
3462	F	427595
3463	F	425701
3464	F	428453
3465	F	426553
3466	F	442272
3467	F	440364
3468	F	443303
3469	F	441380
3470	F	442939
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SEQ ID	Or.	position
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5034	В	717189
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5036	В	718624
5037	В	720503
5038	В	719083
5039	В	720983
5040	В	720047
5041	В	722004
5042	В	720503
5043	В	722393
5044	В	720753
5045	В	722653
5046	В	721798
5047	В	723724
5048	В	722631
5049	В	724493
5050	В	723468
5051	В	725376
5052	В	724852
5053	В	726743
5054	В	726005
5055	В	727903
5056	В	726779
5057	В	728691
5058	В	727058
5059	В	728947
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5061	В	729613
5062	В	728224
5063	В	730116
5064	В	729048
5065	В	730907

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	SEQ ID	Or.	position
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	1877	F	333766
	1878	F	336117
	1879	F	334219
	1880	F	337108
	1881	F	335210
	1882	F	340251
	1883	F	338372
	1884	F	341538
	1885	F	339662
	1886	F	341953
	1887	F	339995
	1888	F	342348
	1889	F	340450
	1890	F	343112
	1891	F	341242
	1892	F	343736
	1893	F	341811
	1894	F	344117
	1895	F	342207
	1896	F	344940
	1897	F	343000
	1898	F	345837
	1899	F	343958
	1900	F	346872
	1901	F	344994
	1902	F	347910
	1903	F	345971
Ī	1904	F	350124
	1905	F	348298
	1906	F	351095
	1907	F	349167
	1908	F	351996
[1909	F	350122

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SEQ ID	Or.	position	
3471	F	441047	
3472	F	445572	
3473	F	443707	
3474	F	467757	
3475	F	465801	
3476	F	471583	
3477	F	469712	
3478	F	487813	
3479	F	485913	
3480	F	496852	
3481	F	494952	
3482	F	499979	
3483	F	498074	
3484	F	508715	
3485	F	506798	
3486	F	510584	
3487	F	508632	
3488	F	526255	
3489	F	524350	
3490	F	531098	
3491	F	529150	
3492	F	556575	
3493	F	554706	
3494	F	564318	
3495	F	562390	
3496	F	566692	
3497	F	564838	
3498	F	570033	
3499	F	568150	
3500	F	570844	
3501	F	568915	
3502	F	575571	
3503	F	573671	
3504	F	590045	

SEQ ID	Or.	position
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5068	В	732909
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5070	В	734663
5071	В	736569
5072		ļ
	В	735879
5073	В	737785
5074	В	736724
5075	В	738632
5076	В	737474
5077	В	739421
5078	В	738007
5079	В	739907
5080	В	738911
5081	В	740799
5082	В	739960
5083	В	741908
5084	В	742277
5085	В	744187
5086	В	743089
5087	В	744989
5088	В	743603
5089	В	745539
5090	В	744565
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5095		747138
5096		745777
5097		747677
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5099	В	748532

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	SEQ ID	Or.	position
	1910	F	353051
	1911	F	351186
	1912	F	353413
	1913	F	351481
	1914	F	353908
	1915	F	351996
	1916	F	354723
	1917	F	352799
	1918	F	356466
İ	1919	F	354569
	1920	F	357107
	1921	F	355178
	1922	F	357 7 67
	1923	F	355878
	1924	F	360528
	1925	F	358628
	1926	F	360877
	1927	F	358974
	1928	F	361573
	1929	F	359692
	1930	F	362584
	1931	F	360681
	1932	F	363835
	1933	F	361966
	1934	F	364960
	1935	F	363021
	1936	F	365240
	1937	F	363360
Ī	1938	F	367060
Ī	1939	F	365115
	1940	F	368383
Ī	1941	F	366505
	1942	F	368862
	1943	F	366963

SEQ ID	Or.	position
3505	F	588196
3506	F	597631
3507	F	595698
3508	F	606387
3509	F	604507
3510	F	607566
3511	F	605637
3512	F	609842
3513	F	607958
3514	F	632472
3515	F	630572
3516	F	636994
3517	F	635071
3518	F	649681
3519	F	647800
3520	F	652059
3521	F	650101
3522	F	654522
3523	F	652562
3524	F	660587
3525	F	658691
3526	F	676785
3527	F	674938
3528	F	679031
3529	F	677133
3530	F	731967
3531	F	730091
3532	F	741797
3533	·F	739935
3534	F	758555
3535	F	756641
3536	F	760010
3537	F	758082
3538	F	770670

SEQ ID	Or.	position
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5101	В	748893
5102	В	748519
5103	В	750396
5104	В	749186
5105	В	751108
5106	В	749646
5107	В	751546
5108	В	749922
5109	В	751824
5110	В	750260
5111	В	752151
5112	В	752527
5113	В	754427
5114	В	753169
5115	В	755064
5116	В	755004
5117	В	756843
5118	В	757034
5119	В	758991
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5121	В	760452
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5123	В	760841
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5125	В	761913
5126	В	760463
5127	В	762363
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5130	В	762053
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5132	В	762363
5133	В	764264

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	SEQ ID	Or.	position
	1944	F	370513
	1945	F	368631
	1946	F	370974
	1947	F	369076
	1948	F	372891
	1949	F	370980
	1950	F	373395
	1951	F	371495
	1952	F	374005
	1953	F	372033
	1954	F	374474
	1955	F	372572
	1956	F	376509
	1957	F	374624
	1958	F	377630
	1959	F	375708
	1960	F	378384
	1961	F	376507
	1962	F	378798
	1963	F	376871
	1964	F	379413
	1965	F	377501
	1966	F	379890
	1967	F	377989
	1968	F	381241
	1969	F	379348
	1970	F	382485
	1971	F	380579
	1972	F	383395
	1973	F	381536
:	1974	F	383730
	1975	F	381782
}	1976	F	384948
	1977	F	383057
Į			

250			
SEQ ID	Or.	position	
3539	F	768751	
3540	F	771896	
3541	F	769996	
3542	F	787857	
3543	F	785958	
3544	F	815714	
3545	F	813840	
3546	F	846380	
3547	F	844470	
3548	F	867576	
3549	F	865673	
3550	F	875167	
3551	F	873254	
3552	F	876214	
3553	F	874314	
3554	F	884093	
3555	F	882162	
3556	F	891248	
3557	F	889348	
3558	F	900125	
3559	F	898298	
3560	F	902048	
3561	F	900125	
3562	F	907563	
3563	F	905656	
3564	F	912076	
3565	F	910133	
3566	F	935157	
3567	F	933211	
3568	F	946473	
3569	F	944568	
3570	F	952562	
3571	F	950664	
3572	F	965649	

SEQ ID	Or.	position
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5135	В	765107
5136	В	764690
5137	В	766595
5138	В	765107
5139	В	766977
5140	В	766327
5141	В	768221
5142	В	766932
5143	В	768851
5144	В	768314
5145	В	770221
5146	В	769045
5147	В	770945
5148	В	770315
5149	В	772234
5150	В	770705
5151	В	772598
5152	В	770882
5153	В	772781
5154	В	771156
5155	В	773044
5156	В	772234
5157	В	774148
5158	В	773611
5159	В	775511
5160	В	774513
5161	В	776404
5162	В	776333
5163	В	778191
5164	В	777926
5165	В	779832
5166	В	777455
5167	В	779380

SEQ ID	Or.	position
1978	F	385474
1979	F	383532
1980	F	385908
1981	F	384008
1982	F	386643
1983	F	384750
1984	F	387099
1985	F	385204
1986	F	387581
1987	F	385677
1988	F	388009
1989	F	386062
1990	F	388927
1991	F	387033
1992	F	389726
1993	F	387821
1994	F	391295
1995	F	389365
1996	F	392171
1997	F	390291
1998	F	393930
1999	F	392014
2000	F	395085
2001	F	393185
2002	F	395827
2003	F	393940
2004	F	396274
2005	F	394423
2006	F	397156
2007	F	395216
2008	F	398641
2009	F	396790
2010	F	399550
2011	F	397659

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SEQ ID	Or.	position
3573	F	963730
3574	F	968519
3575	F	966614
3576	F	970497
3577	F	968601
3578	F	971879
3579	F	970043
3580	F	972888
3581	F	970962
3582	F	998162
3583	F	996241
3584	F	1003657
3585	F	1001756
3586	F	1009313
3587	F	1007413
3588	F	1028954
3589	F	1027039
3590	В	730
3591	В	2645
3592	В	3521
3593	В	5431
3594	В	5295
3595	В	7188
3596	В	6740
3597	В	8652
3598	В	8240
3599	В	10138
3600	В	8959
3601	В	10816
3602	В	9285
3603	В	11160
3604	В	9689
3605	В	11591
3606	В	10679

SEQ ID	Or.	position
5168	B	779476
5169	В	781342
5170	В	781774
5171	В	783686
5172	В	782667
5173	В	784562
5174	В	785748
5175	В	787658
5176	В	786222
5177	В	788126
5178	В	786803
5179	В	788703
5180	В	787998
5181	В	789876
5182	В	788279
5183	В	790255
5184	В	790369
5185	В	792247
5186	В	790862
5187	В	792787
5188	В	792247
5189	В	794137
5190	В	793352
5191	В	795215
5192	В	794276
5193	В	796196
5194	В	795215
5195	В	797077
5196	В	795667
5197	В	797571
5198	В	796515
5199	В	798382
5200	В	797235
5201	В	799168

SEQ ID	Or.	position
2012	F	399797
2013	F	397915
2014	F	401527
2015	F	399623
2016	F	401907
2017	F	399960
2018	F	403017
2019	F	401131
2020	F	403017
2021	F	401131
2022	F	404910
2023	F	403010
2024	F	405728
2025	F	403836
2026	F	406837
2027	F	404932
2028	F	410291
2029	F	408347
2030	F	411488
2031	F	409518
2032	F	412379
2033	F	410487
2034	F	413164
2035	F	411263
2036	F	413606
2037	F	411626
2038	F	413721
2039	F	411859
2040	F	414921
2041	F	413049
2042	F	416517
2043	F	414606
2044	F	417445
2045	F	415557

SEQ	ID	Or.	position
360	7	В	12568
360	8	В	11515
360	9	В	13376
361	0	В	12136
361	1	В	14038
361	2	В	13565
361	3	В	15450
361	4	В	14667
361	5	В	16535
361	6	В	15254
361	7	В	17183
361	В	В	16189
361	9	В	18097
362	0	В	16790
362	i	В	18621
362	2	В	16710
362	3	В	18587
3624	1	В	18458
362	5	В	20318
3626	5	В	20768
362	7	В	22647
3628	3	В	22438
3629	•	В	24335
3630)	В	22892
3631	-	В	24752
3632	2	В	23046
3633	3	В	24924
3634		В	23442
3635	5	В	25313
3636		В	23623
3637	<u>'</u>	В	25528
3638		В	24822
3639)	В	26708
3640		В	26618
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SEQ ID	Or.	<u> </u>
5202	В	797585
5203	В	799553
5204	В	798102
5205	В	799976
5206	В	798306
5207	В	800214
5208	В	798662
5209	В	800546
5210	В	799168
5211	В	801074
5212	В	800159
5213	В	802054
5214	В	801153
5215	В	803031
5216	В	802363
5217	В	804257
5218	В	802893
5219	В	804802
5220	В	803466
5221	В	805366
5222	В	804440
5223	В	806374
5224	В	805576
5225	В	807523
5226	В	806511
5227	В	808410
5228		808305
5229		810207
5230		809229
5231	i	811164
5232	1	810700
5233		812581
5234		811825
5235	В	813787

SEQ ID	Or.	position
2046	F	417812
2047	F	415912
2048	F	418381
2049	F	416517
2050	F	419453
2051	F	417447
2052	F	420026
2053	F	418068
2054	F	421270
2055	F	419392
2056	F	422262
2057	F	420364
2058	F	423007
2059	F	421107
2060	F	424834
2061	F	423018
2062	F	426026
2063	F	424110
2064	F	426883
2065	F	425040
2066	F	429303
2067	F	427438
2068	F	429609
2069	F	427737
2070	F	430104
2071	F	428191
2072	F	430622
2073	F	428814
2074	F	431439
2075	F	429537
2076	F	431968
2077	F	430108
2078	F	434401
2079	F	432485

3641 3642	Or. B	28544
3642	В	
	 	27072
	1	27972
3643	B	29859
3644	В	29000
3645	В	30908
3646	В	30242
3647	В	32160
3648	В	30966
3649	В	32849
3650	В	31516
3651	В	33401
3652	В	32032
3653	В	33900
3654	В	33576
3655	В	35496
3656	В	34121
3657	В	36052
3658	В	35251
3659	В	37164
3660	В	35588
3661	В	37502
3662	В	36152
3663	В	38041
3664	В	37216
3665	В	39127
3666	В	38041
3667	В	39980
3668	В	39322
3669	В	41205
3670	В	40161
3671	В	42074
3672	В	40985
3673	В	42887
3674	В	42284

SEQ ID	Or.	position	
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5239	В	815676	
5240	В	814082	
5241	В	815892	
5242	В	814571	
5243	В	816454	
5244	В	815441	
5245	В	817347	
5246	В	815101	
5247	В	817025	
5248	В	815953	
5249	В	817887	
5250	В	817709	
5251	В	819618	
5252	В	819559	
5253	В	821442	
5254	В	819933	
5255	В	821846	
5256	В	820622	
5257	В	822548	
5258	В	821281	
5259	В	823181	
5260	В	821778	
5261	В	823666	
5262	В	822494	
5263	В	824387	
5264	В	823155	
5265	В	825057	
5266	В	824589	
5267	В	826509	
5268	В	826122	
5269	В	828041	

	WO 99/28475		
	SEQ ID	Or.	position
	2080	F	436050
	2081	F	434134
	2082	F	436342
	2083	F	434428
	2084	F	437672
	2085	F	435775
	2086	F	438051 -
	2087	F	436165
	2088	F	438767
	2089	F	436866
	2090	F	439139
	2091	F	437145
	2092	F	439479
	2093	F	437574
	2094	F	440823
	2095	F	438923
	2096	F	441668
	2097	F	439746
	2098	F	444271
	2099	F	442371
	2100	F	446233
·	2101	F	444302
	2102	F	447687
	2103	F	445803
	2104	F	450318
	2105	F	448399
	2106	F	450876
	2107	F	449025
	2108	F	451274
	2109	F	449397
	2110	F	452413
	2111	F	450513
	2112	F	453303
	2113	F	451427

SEQ ID	Or.	position	
3675	В	44180	
3676	В	43337	
3677	В	45275	
3678	В	44022	
3679	В	45927	
3680	В	45451	
3681	В	47375	
3682	В	46308	
3683	В	48208	
3684	В	46560	
3685	В	48485	
3686	В	47115	
3687	В	49053	
3688	В	48208	
3689	В	50118	
3690	В	48806	
3691	В	50708	
3692	В	50333	
3693	В	52220	
3694	В	50960	
3695	В	52890	
3696	В	52660	
3697	В	54606	
3698	В	53737	
3699	В	55645	
3700	В	54793	
3701	В	56691	
3702	В	55329	
3703	В	57226	
3704	В	56691	
3705	В	58566	
3706	В	56317	
3707	В	58226	
3708	В	58469	

SEQ ID	Or.	position
5270	В	826572
5271	В	828500
5272	В	827246
5273	В	829208
5274	В	827489
5275	В	829407
5276	В	828274
5277	В	830176
5278	В	829495
5279	В	831395
5280	В	831192
5281	В	833082
5282	В	832005
5283	В	833870
5284	В	832373
5285	В	834280
5286	В	832964
5287	В	834864
5288	В	833672
5289	В	835604
5290	В	834022
5291	В	835945
5292	В	834907
5293	В	836803
5294	В	835683
5295	В	837593
5296	В	836018
5297	В	837902
5298	В	836714
5299	В	838602
5300	В	837484
5301	В	839424
5302	В	838950
5303	В	840832

SEQ ID	Or.	position
2114	F	454713
2115	F	452853
2116	F	455096
2117	F	453248
2118	F	455808
2119	F	453928
2120	F	457440
2121	F	455488
2122	F	458282
2123	F	456354
2124	F	459558
2125	F	457686
2126	F	460960
2127	F	459060
2128	F	461659
2129	F	459758
2130	F	462674
2131	F	460775
2132	F	463788
2133	F	461895
2134	F	464479
2135	F	462602
2136	F	465882
2137	F	463989
2138	F	467200
2139	F	465300
2140	F	468680
2141	F	466787
2142	F	469130
2143	F	467224
2144	F	469572
2145	F	467707
2146	F	470887
2147	F	468984

SEQ ID	Or.	position	
3709	В	60363	
3710	В	59406	
3711	В	61301	
3712	В	60450	
3713	В	62316	
3714	В	61722	
3715	В	63603	
3716	В	62585	
3717	В	64473	
3718	В	63362	
3719	В	65285	
3720	В	64203	
3721	В	66103	
3722	В	64850	
3723	В	66749	
3724	В	64899	
3725	В	66776	
3726	В	65164	
3727	В	67067	
3728	В	67514	
3729	В	69390	
3730	В	69097	
3731	В	71033	
3732	В	69470	
3733	В	71354	
3734	В	69961	
3735	В	71898	
3736	В	70707	
3737	В	72579	
3738	В	71249	
3739	В	73134	
3740	В	73458	
3741	В	75376	
3742	В	75121	

Or.	position
В	839800
В	841700
В	840808
В	842715
В	841975
В	843834
В	842679
В	844614
В	843342
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В	843736
В	845626
В	846423
В	848330
В	844423
В	846258
В	848265
В	850174
В	850343
В	852246
В	851875
L 1	853765
В	853944
В	855822
В	855056
В	856964
В	855750
В	857714
В	856488
В	858370
В	857403
В	859303
В	859659
В	861570
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•	WO 99/28475		
	SEQ ID	Or.	position
	2148	F	471590
	2149	F	469835
	2150	F	473033
	2151	F	471133
	2152	F	473761
i	2153	F	471861
	2154	F	474383
	2155	F	472478
	2156	F	475333
	2157	F	473465
	2158	F	476279
	2159	F	474390
	2160	F	478446
	2161	F	476546
	2162	F	478869
	2163	F	476918
Ī	2164	F	479441
	2165	F	477548
	2166	F	479676
	2167	F	477775
Ī	2168	F	481277
	2169	F	479377
	2170	F	481635
	2171	F	479745
	2172	F	483172
	2173	F	481279
Ī	2174	F	484659
	2175	F	482764
Ţ	2176	F	485003
Ī	2177	F	483097
	2178	F	487946
	2179	F	486083
	2180	F	487946
	2181	F	486093

SEQ ID	Or.	position	
3743	В	77020	
3744	В	75741	
3745	В	77608	
3746	В	77447	
3747	В	79300	
3748	В	77535	
3749	В	79430	
3750	В	78333	
3751	В	80243	
3752	В	79300	
3753	В	81197	
3754	В	82201	
3755	В	84100	
3756	В	83493	
3757	В	85405	
3758	В	85075	
3759	В	86973	
3760	В	87141	
3761	В	89086	
3762	В	88903	
3763	В	90810	
3764	В	88038	
3765	В	89946	
3766	В	88981	
3767	В	90884	
3768	В	89485	
3769	В	91385	
3770	В	89975	
3771	В	91855	
3772	В	91680	
3773	В	93567	
3774	В	92008	
3775	В	93902	
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SEQ ID	Or.	position
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5345	В	864173
5346	В	863792
5347	В	865660
5348	В	864164
5349	В	866050
5350	В	864925
5351	В	866860
5352	В	866468
5353	В	868368
5354	В	866860
5355	В	868734
5356	В	867952
5357	В	869878
5358	В	869311
5359	В	871226
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5361	В	871517
5362	В	871614
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5365	В	874702
5366	В	873093
5367	В	874980
5368	В	874158
5369	В	876056
5370	В	874033
5371	В	875932

SEQ ID Or. position 2182 F 489220 2183 F 487274 2184 F 490276 2185 F 488388 2186 F 492138 2187 F 490229 2188 F 490475 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 495569 2204 F 498050 2	 	WO 99/28475		
2183 F 487274 2184 F 490276 2185 F 488388 2186 F 492138 2187 F 490229 2188 F 490229 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498671 2210 <th></th> <th>SEQ ID</th> <th>Or.</th> <th>position</th>		SEQ ID	Or.	position
2184 F 490276 2185 F 488388 2186 F 492138 2187 F 490229 2188 F 490475 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 498050 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2210 F 502469 <t< td=""><td></td><td>2182</td><td>F</td><td>489220</td></t<>		2182	F	489220
2185 F 488388 2186 F 492138 2187 F 490229 2188 F 4902475 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 498569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2210 F 501547 2211 F 502469 <		2183	F	487274
2186 F 492138 2187 F 490229 2188 F 490475 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2210 F 501547 2211 F 502469 2213 F 503435 <td></td> <td>2184</td> <td>F</td> <td>490276</td>		2184	F	490276
2187 F 490229 2188 F 492475 2189 F 490618 2190 F 493591 2191 F 491719 2192 F 494297 2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 497468 2202 F 497468 2203 F 498050 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2210 F 501547 2211 F 502469 2212 F 502469 2213 F 503435 <td></td> <td>2185</td> <td>F</td> <td>488388</td>		2185	F	488388
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2193 F 492436 2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2191	F	491719
2194 F 494530 2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2192	F	494297
2195 F 492679 2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2193	F	492436
2196 F 494637 2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2194	F	494530
2197 F 492753 2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2195	F	492679
2198 F 495467 2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2196	F	494637
2199 F 493559 2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2197	F	492753
2200 F 496076 2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2198	F	495467
2201 F 494126 2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2199	F	493559
2202 F 497468 2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2200	F	496076
2203 F 495569 2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2201	F	494126
2204 F 498050 2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2202	F	497468
2205 F 496160 2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2203	F	495569
2206 F 498994 2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2204	F	498050
2207 F 497096 2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2205	F	496160
2208 F 500571 2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2206	F	498994
2209 F 498671 2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2207	F	497096
2210 F 501547 2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2208	F	500571
2211 F 499671 2212 F 502469 2213 F 500572 2214 F 503435		2209	F	498671
2212 F 502469 2213 F 500572 2214 F 503435		2210	F	501547
2213 F 500572 2214 F 503435		2211	F	499671
2214 F 503435		2212	F	502469
	•	2213	F	500572
2215 F 501547		2214	F	503435
		2215	F	501547

201			
SEQ ID	Or.	position	
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3779	В	95288	
3780	В	94354	
3781	В	96254	
3782	В	94897	
3783	В	96772	
3784	В	98519	
3785	В	100439	
3786	В	98962	
3787	В	100853	
3788	В	100451	
3789	В	102387	
3790	В	101639	
3791	В	103473	
3792	В	102457	
3793	В	104357	
3794	В	102745	
3795	В	104666	
3796	В	104544	
3797	В	106464	
3798	В	105338	
3799	В	107237	
3800	В	106127	
3801	В	108014	
3802	В	108693	
3803	В	110587	
3804	В	109189	
3805	В	111079	
3806	В	109684	
3807	В	111580	
3808	В :	110587	
3809	В :	112486	
3810	В :	112740	

SEQ ID	Or.	position
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5374	В	875606
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5376	В	876619
5377	В	878519
5378	В	878160
5379	В	880034
5380	В	878429
5381	В	880292
5382	В	879336
5383	В	881230
5384	В	879393
5385	В	881281
5386	В	880089
5387	В	881994
5388	В	881108
5389	В	883062
5390	В	882094
5391	В	884013
5392	В	883124
5393	В	885027
5394	В	884014
5395	В	885924
5396	В	884530
5397	В	886432
5398	В	885226
5399	В	887139
5400	В	889010
5401	В	890900
5402	В	890421
5403	В	892310
5404	В	891438
5405	В	893287
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SEQ ID	Or.	position
2216	F	505469
2217	F	503545
2218	F	506768
2219	F	504880
2220	F	507356
2221	F	505530
2222	F	508015
2223	F	506157
2224	F	508247
2225	F	506351
2226	F	509270
2227	F	507356
2228	F	510759
2229	F	508918
2230	F	511268
2231	F	509359
2232	F	512124
2233	F	510202
2234	F	512836
2235	F	510926
2236	F	514569
2237	F	512663
2238	F	514688
2239	F	512874
2240	F	515334
2241	F	513415
2242	F	515516
2243	F	513670
2244	F	516230
2245	F	514348
2246	F	517181
2247	F	515261
2248	F	517840
2249	F	515977

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SEQ ID	Or.	
3811	В	114673
3812	В	113684
3813	В	115653
3814	В	114216
3815	В	116158
3816	В	114836
3817	В	116732
3818	В	115473
3819	В	117380
3820	В	115898
3821	В	117797
3822	В	120031
3823	В	121926
3824	В	124231
3825	В	126158
3826	В	125215
3827	В	127115
3828	В	125352
3829	В	127271
3830	В	126492
3831	В	128390
3832	В	127150
3833	В	129050
3834	В	128010
3835	В	129905
3836	В	129947
3837	В	131873
3838	В	131255
3839	В	133205
3840	В	131873
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3842	В	133381
3843	В	135305
3844	В	133804

SEQ ID	Or.	position
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5407	В	893606
5408	В	893606
5409	В	895490
5410	В	894049
5411	B	896024
5412	В	894139
5413	В	896074
5414	В	894545
5415	·B	896413
5416	В	894999
5417	В	896912
5418	В	896127
5419	В	898012
5420	В	897049
5421	В	898949
5422	В	901018
5423	В	902955
5424	В	902393
5425	В	904301
5426	В	904098
5427	В	906002
5428	В	903951
5429	В	905851
5430	В	905825
5431	В	907725
5432	В	906700
5433	В	908669
5434	В	907174
5435	В	909066
5436	В	907579
5437	В	909480
5438	В	908962
5439	В	910922

SEQ ID	Or.	position
2250	F	518184
2251	F	516283
2252	F	518704
2253	F	516804
2254	F	519358
2255	F	517404
2256	F	520483
2257	F	518598
2258	F	521632
2259	F	519765
2260	F	521919
2261	F	520024
2262	F	523372
2263	F	521495
2264	F	524404
2265	F	522508
2266	F	525498
2267	F	523600
2268	F	525687
2269	F	523753
2270	F	526036
2271	F	524164
2272	F	526632
2273	F	524732
2274	F	528073
2275	F	526158
2276	F	528835
2277	F	526999
2278	F	529393
2279	F	527504
2280	F	530004
2281	F	528082
2282	F	530518
2283	F	528594

SEQ ID	Or.	position
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3846	В	134219
3847	В	136133
3848	В	134729
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3850	В	137694
3851	В	139583
3852	В	140496
3853	В	142396
3854	В	140953
3855	В	142856
3856	В	142031
3857	В	143950
3858	В	143520
3859	В	145425
3860	В	144066
3861	В	145967
3862	В	144629
3863	В	146519
3864	В	146547
3865	В	148446
3866	В	147304
3867	В	149227
3868	В	148296
3869	В	150255
3870	В	148769
3871	В	150670
3872	В	149320
3873	В	151168
3874	В	149532
3875	В	151470
3876	В	149934
3877	В	151845
3878	В	151168

SEQ ID	Or.	position
5440	В	909415
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5442	В	909804
5443	В	911704
5444	B	910218
5445	В	912075
5446	В	910834
5447	В	912689
5448	В	911225
5449	В	913064
5450	В	912492
5451	В	914381
5452	В	912852
5453	В	914763
5454	В	913438
5455	В	915340
5456	В	913901
5457	В	915814
5458	В	914730
5459	В	916694
5460	В	915356
5461	В	917256
5462	В	915717
5463	В	917608
5464	В	916854
5465	В	918761
5466	В	916762
5467	В	918712
5468	В	917848
5469	В	919756
5470	В	918588
5471	В	920424
5472	В	918875
5473	В	920796
L		

SEQ ID Or. position 2284 F 532100 2285 F 530224 2286 F 533025 2287 F 531125 2288 F 533710 2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 53877 2298 F 540672 2299 F 538777 2300 F 541573 2301 F 539706		WO 99/28475			
2285 F 530224 2286 F 533025 2287 F 531125 2288 F 533710 2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		position	Or.	SEQ ID	
2286 F 533025 2287 F 531125 2288 F 533710 2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		532100	F	2284	
2287 F 531125 2288 F 533710 2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573	_	530224	F	2285	
2288 F 533710 2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		533025	F	2286	
2289 F 531825 2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		531125	F	2287	
2290 F 534658 2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		533710	F	2288	
2291 F 532758 2292 F 535737 2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		531825	F	2289	
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2293 F 533828 2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		532758	F	2291	
2294 F 539456 2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		535737	F	2292	
2295 F 537568 2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573		533828	F	2293	
2296 F 540290 2297 F 538375 2298 F 540672 2299 F 538777 2300 F 541573	_	539456	F	2294	
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2298 F 540672 2299 F 538777 2300 F 541573		540290	F	2296	:
2299 F 538777 2300 F 541573		538375	F	2297	
2300 F 541573	_	540672	F	2298	
		538777	F	2299	
2301 F 539706		541573	F	2300	
		539706	F	2301	
2302 F 542999		542999	F	2302	
2303 F 541102		541102	F	2303	
2304 F 543922		543922	F	2304	Ì
2305 F 542057		542057	F	2305	
2306 F 544268		544268	F	2306	
2307 F 542354		542354	F	2307	
2308 F 544691	٦	544691	F	2308	`
2309 F 542785	┨	542785	F	2309	
2310 F 546323	٦	546323	F	2310	
2311 F 544441	٦	544441	F	2311	
2312 F 546467	٦	546467	F	2312	
2313 F 544650	\dashv	544650	F	2313	
2314 F 547257	\dashv	547257	F	2314	Ī
2315 F 545357	\dashv	545357	F	2315	
2316 F 547658	\dashv	547658	F	2316	ſ
2317 F 545781	7	545781	F	2317	

200			
SEQ ID	Or.	position	
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3881	В	154165	
3882	В	152887	
3883	В	154804	
3884	В	153600	
3885	В	155482	
3886	В	154217	
3887	В	156157	
3888	В	155523	
3889	В	157396	
3890	В	156823	
3891	В	158731	
3892	В	157745	
3893	В	159645	
3894	В	159207	
3895	В	161084	
3896	В	160216	
3897	В	162127	
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3899	В	163350	
3900	В	161818	
3901	В	163726	
3902	В	162601	
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3904	В	163282	
3905	В	165205	
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3908	В	165812	
3909	В	167672	
3910	В	166939	
3911	В	168861	
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SEQ ID	Or.	position
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5476	В	920055
5477	В	921949
5478	В	920389
5479	В	922328
5480	В	921130
5481	В	922978
5482	В	921517
5483	В	923414
5484	В	921740
5485	В	923646
5486	В	921979
5487	В	923926
5488	В	922396
5489	В	924327
5490	В	922729
5491	В	924611
5492	В	923256
5493	В	925216
5494	В	923673
5495	В	925589
5496	В	924297
5497	В	926176
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5500	В	925216
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5504	В	926980
5505	В	928903
5506	В	927408
5507	В	929331

SEQ ID	Or.	position
2318	F	548406
2319	F	546466
2320	F	549443
2321	F	547527
2322	F	550193
2323	F	548288
2324	F	551554
2325	F	549665
2326	F	552877
2327	F	550963
2328	F	554708
2329	F	552807
2330	F	556293
2331	F	554381
2332	F	557095
2333	F	555164
2334	F	557522
2335	F	555656
2336	F	558021
2337	F	556191
2338	F	558964
2339	F	557076
2340	F	560913
2341	F	559007
2342	F	562832
2343	F	561002
2344	F	563374
2345	F	561474
2346	F	565061
2347	F	563198
2348	F	566840
2349	F	564908
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2351	F	565896

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SEQ ID	Or.	position
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2429	F	602388
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2433	F	604721
2434	F	607722
2435	F	605908
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2438	F	609842
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2440	F	611565
2441	F	609657
2442	F	612719
2443	F	610819
2444	F	613473
2445	F	611565
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SEQ ID	Or.	position
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4041	В	227907
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4045	В	228871
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5637	В	990252
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5640	В	989437
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SEQ ID	Or.	position
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5664	В	1004003
5665	В	1005872
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SEQ ID	Or.	position
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2512	F	651421
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4112	В	266644
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WHAT IS CLAIMED IS:

- 1. An isolated polynucleotide having a nucleotide sequence of a *Chlamydia trachomatis* genome, comprising
 - (a) the nucleotide sequence of SEQ ID No. 1;
 - the nucleotide sequence contained within the Chlamydia trachomatis genomic
 DNA in ECACC Deposit No. 98112618;
 - (c) the nucleotide sequence contained in a clone insert in ECACC Deposit No. 98112617;
 - (d) a nucleotide sequence exhibiting at least 99.9% identity with the sequence of SEQ ID No. 1; or
 - (e) a nucleotide sequence exhibiting at least 80% homology to SEQ ID No. 1.
- 2. An isolated polynucleotide which hybridizes to SEQ ID No. 1 or to the Chlamydia trachomatis genomic DNA contained in ECACC Deposit No. 98112618 or to a clone insert in ECACC Deposit No. 98112617 under conditions of high stringency.
- 3. An isolated polynucleotide which hybridizes to SEQ ID No. 1 or to the Chlamydia trachomatis genomic DNA contained in ECACC Deposit No. 98112618 under conditions of intermediate stringency.
- 4. An isolated polynucleotide having a nucleotide sequence of an open reading frame (ORF) of a *Chlamydia trachomatis* genome, comprising:
 - (a) a nucleotide sequence chosen from one of ORF2 to ORF 1197;
 - (b) a nucleotide sequence exhibiting at least 99.9% identity with one of ORF2 to ORF 1197; or
 - (c) a nucleotide sequence exhibiting at least 80% homology to one of ORF2 to ORF 1197.
- 5. An isolated polynucleotide which hybridizes to one of ORF2 to ORF 1197 under conditions of high stringency.
- 6. An isolated polynucleotide which hybridizes to one of ORF2 to ORF 1197 under conditions of intermediate stringency.
- 7. The polynucleotide of Claim 2, 3, 4, 5, or 6 which encodes the following polypeptides or fragments thereof:

- (a) a Chlamydia trachomatis transmembrane polypeptide having between 1 and 3 transmembrane domains;
- (b) a *Chlamydia trachomatis* transmembrane polypeptide having between 4 and 6 transmembrane domains;
- (c) a *Chlamydia trachomatis* transmembrane polypeptide having at least 7 transmembrane domains;
- (d) a *Chlamydia trachomatis* polypeptide involved in intermediate metabolism of sugars and/or cofactors;
- (e) a Chlamydia trachomatis polypeptide involved in intermediate metabolism of nucleotides or nucleic acids;
- (f) a Chlamydia trachomatis polypeptide involved in metabolism of amino acids or polypeptides;
- (g) a Chlamydia trachomatis polypeptide having involved in metabolism of fatty acids;
- (h) a Chlamydia trachomatis polypeptide involved in the synthesis of the cell wall;
- a Chlamydia trachomatis polypeptide involved in transcription, translation, and/or maturation process;
- (j) a Chlamydia trachomatis transport polypeptide:
- (k) a Chlamydia trachomatis polypeptide involved in the virulence process;
- a Chlamydia trachomatis polypeptide involved in the secretory system and/or which is secreted;
- (m) a Chlamydia trachomatis polypeptide of the cellular envelope or outer cellular envelope of Chlamydia trachomatis.
- (n) a Chlamydia trachomatis surface exposed polypeptide;
- (o) a Chlamydia trachomatis lipoprotein;
- a Chlamydia trachomatis polypeptide involved in lipopolysaccharide biosynthesis;
- (q) a Chlamydia trachomatis KDO-related polypeptide;
- (r) a Chlamydia trachomatis phosphomannomutase-related polypeptide;
- (s) a Chlamydia trachomatis phosphoglucomutase-related polypeptide;
- (t) a Chlamydia trachomatis lipid A component-related polypeptide;
- (u) a Chlamydia trachomatis polypeptide that contains an RGD sequence;
- (v) a Chlamydia trachomatis Type III secreted polypeptide;
- (w) a Chlamydia trachomatis cell wall anchored surface polypeptide; or
- (x) a Chlamydia trachomatis polypeptide that is not found in Chlamydia trachomatis.

- 8. A polynucleotide encoding a fusion protein, comprising one of ORF2 to ORF 1197 of Claim 4, 5, or 6 ligated in frame to a polynucleotide encoding a heterologous polypeptide.
- 9. A recombinant vector that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
 - 10. A recombinant vector that contains the polynucleotide of Claim 8.
- 11. A recombinant vector that contains the polynucleotide of Claim 4, 5 or 6, operatively associated with a regulatory sequence that controls gene expression.
- 12. A recombinant vector that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression.
- 13. A genetically engineered host cell that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
 - 14. A genetically engineered host cell that contains the polynucleotide of Claim 8.
- 15. A genetically engineered host cell that contains the polynucleotide of Claim 4, 5 or 6 operatively associated with a regulatory sequence that controls gene expression in the host cell.
- 16. A genetically engineered host cell that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression in the host cell.
 - 17. A method for producing a polypeptide, comprising:
 - (a) culturing the genetically engineered host cell of Claim 15 under conditions suitable to produce the polypeptide encoded by the polynucleotide; and
 - (b) recovering the polypeptide from the culture.
 - 18. A method for producing a fusion protein, comprising:
 - (a) culturing the genetically engineered host cell of Claim 16 under conditions suitable to produce the fusion protein encoded by the polynucleotide; and
 - (b) recovering the fusion protein from the culture.
 - 19. A polypeptide encoded by the polynucleotide of Claim 4, 5 or 6.

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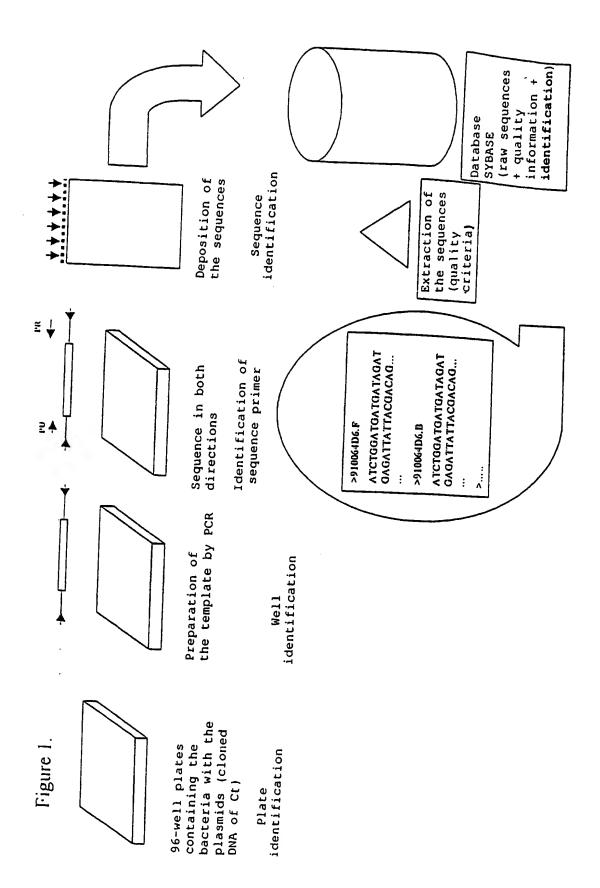
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- 20. The polypeptide of Claim 19 which immunoreacts with seropositive serum of an individual infected with *Chlamydia trachomatis*.
- 21. The polypeptide of Claim 19 which comprises the following polypeptides or fragments thereof:
 - (a) a Chlamydia trachomatis transmembrane polypeptide having between 1 and 3 transmembrane domains;
 - (b) a *Chlamydia trachomatis* transmembrane polypeptide having between 4 and 6 transmembrane domains;
 - (c) a *Chlamydia trachomatis* transmembrane polypeptide having at least 7 transmembrane domains;
 - (d) a Chlamydia trachomatis polypeptide involved in intermediate metabolism of sugars and/or cofactors;
 - (e) a Chlamydia trachomatis polypeptide involved in intermediate metabolism of nucleotides or nucleic acids;
 - a Chlamydia trachomatis polypeptide involved in metabolism of amino acids or polypeptides;
 - (g) a Chlamydia trachomatis polypeptide involved in metabolism of fatty acids;
 - (h) a Chlamydia trachomatis polypeptide involved in the synthesis of the cell wall;
 - a Chlamydia trachomatis polypeptide involved in transcription, translation, and/or maturation process;
 - (j) a Chlamydia trachomatis transport polypeptide;
 - (k) a Chlamydia trachomatis polypeptide involved in the virulence process;
 - (l) a Chlamydia trachomatis polypeptide involved in the secretory system and/or which is secreted;
 - (m) a Chlamydia trachomatis polypeptide of the cellular envelope or outer cellular envelope of Chlamydia trachomatis.
 - (n) a Chlamydia trachomatis surface exposed polypeptide;
 - (o) a Chlamydia trachomatis lipoprotein;
 - a Chlamydia trachomatis polypeptide involved in lipopolysaccharide biosynthesis;
 - (q) a Chlamydia trachomatis KDO-related polypeptide;
 - (r) a Chlamydia trachomatis phosphomannomutase-related polypeptide;
 - (s) a Chlamydia trachomatis phosphoglucomutase-related polypeptide;
 - (t) a Chlamydia trachomatis lipid A component-related polypeptide;

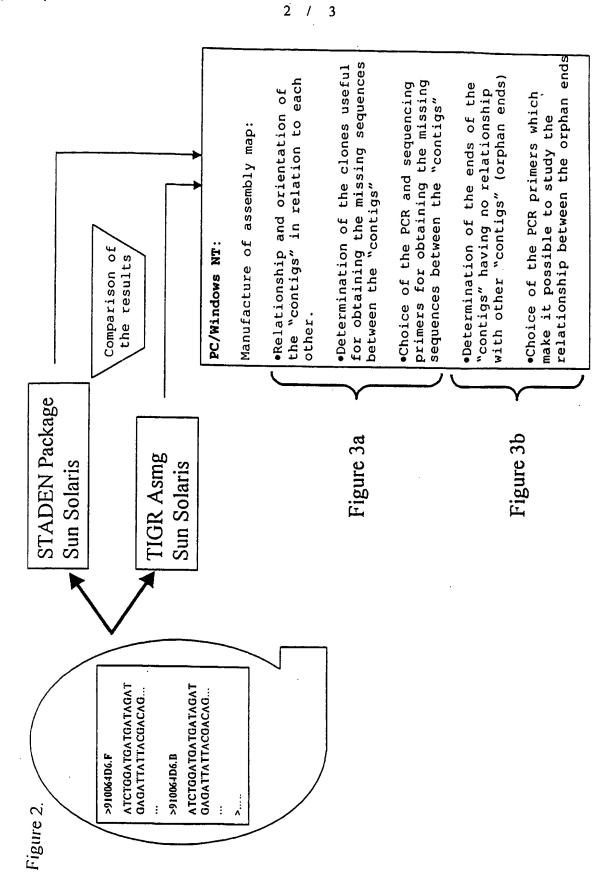
- (u) a Chlamydia trachomatis polypeptide that contains an RGD sequence;
- (v) a Chlamydia trachomatis Type III secreted polypeptide;
- (w) a Chlamydia trachomatis cell wall anchored surface polypeptide; or
- (x) a Chlamydia trachomatis polypeptide that is not found in Chlamydia trachomatis.
- 22. A fusion protein encoded by the polynucleotide of Claim 8.
- 23. The fusion protein of Claim 22 which immunoreacts with seropositive serum of an individual infected with *Chlamydia trachomatis*.
 - 24. An antibody that immunospecifically binds to the polypeptide of Claim 19.
 - 25. An antibody that immunospecifically binds to the fusion protein of Claim 22.
- 26. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
 - (a) contacting the sample with a polynucleotide primer of Claim 1, 2, 3, 4, 5, or 6 in the presence of a polymerase enzyme and nucleotides under conditions which permit primer extension; and
 - (b) detecting the presence of primer extension products in the sample in which the detection of primer extension products indicates the presence of *Chlamydia trachomatis* in the sample.
- 27. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
 - (a) contacting the sample with a polynucleotide probe of Claim 1, 2, 3, 4, 5, or 6 under conditions which permit hybridization of complementary base pairs; and
 - (b) detecting the presence of hybridization complexes in the sample in which the detection of hybridization complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 28. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
 - (a) contacting the sample with the antibody of Claim 24 under conditions suitable for the formation of immune complexes; and

- (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 29. A method for the detection and/or identification of antibodies to *Chlamydia trachomatis* in a biological sample, comprising:
 - (a) contacting the sample with a polypeptide of Claim 19 under conditions suitable for the formation of immune complexes; and
 - (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 30. A DNA chip containing an array of polynucleotides comprising at least one of the polynucleotides of Claim 1, 2, 3, 4, 5, or 6.
- 31. A protein chip containing an array of polypeptides comprising at least one of the polypeptides of Claim 19.
- 32. An immunogenic composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 33. An immunogeneic composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.
- 34. An immunogenic composition comprising the fusion protein of Claim 22 and a pharmaceutically acceptable carrier.
- 35. An immunogenic composition comprising the fusion protein of Claim 23 and a pharmaceutically acceptable carrier.
- 36. A pharmaceutical composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 37. A pharmaceutical composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.

- 38. A pharmaceutical composition comprising the polypeptide of Claim 22 and a pharmaceutically acceptable carrier.
- 39. A pharmaceutical composition comprising the polypeptide of Claim 23 and a pharmaceutically acceptable carrier.
- 40. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 32.
- 41. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 33.
- 42. A method of immunizing against *Chlamydia trachomatis*, comprising administering to a host an immunizing amount of the immunogenic composition of Claim 34.
- 43. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 35.
- 44. A DNA immunogenic composition comprising the expression vector of Claim 11.
- 45. The DNA composition of Claim 44, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia trachomatis*.
- 46. A DNA immunogenic composition comprising the expression vector of Claim 12.
- 47. The DNA composition of Claim 46, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia trachomatis*.
 - 48. A screening assay, comprising:
 - (a) contacting a test compound with an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6; and
 - (b) detecting whether binding occurs.
 - 49. A screening assay, comprising:
 - (a) contacting a test compound with the polypeptide of Claim 19; and

- (b) detecting whether binding occurs.
- 50. A screening assay, comprising:
- (a) contacting a test compound with the polypeptide of Claim 22; and
- (b) detecting whether binding occurs.
- 51. A kit comprising a container containing an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
 - 52. The kit of Claim 51 wherein the polynucleotide is a primer or a probe.
- 53. The kit of Claim 51 wherein the polynucleotide is a primer and the kit further comprises a container containing a polymerase.
- 54. The kit of Claim 51 which further comprises a container containing deoxynucleotide triphosphates.
- 55. A kit comprising a container containing an antibody that immunospecifically binds to the polypeptide of Claim 19.
- 56. A kit comprising a container containing an antibody that immunospecifically binds to the fusion protein of Claim 22.





2- Define the PCR primers which make it possible to fill

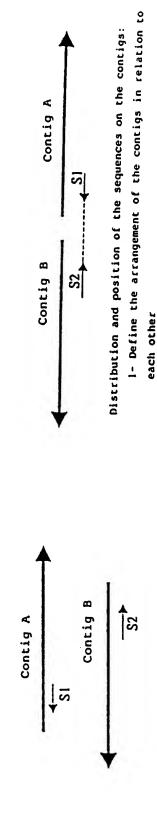
the sequence

Statistical determination of the sequences:

1- Belonging to the same clone

2- Situated on two different contigs

FIGURE 3A



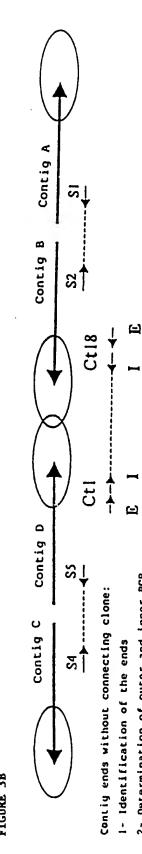


FIGURE 3B

E: outer primers

between the contigs

2- Determination of outer and inner PCR primers for studying the relationships

1: inner primers